

10/009,574

=> d his

(FILE 'HOME' ENTERED AT 15:55:36 ON 11 JUN 2003)

FILE 'REGISTRY' ENTERED AT 15:55:41 ON 11 JUN 2003

L1 1 S QUETIAPINE/CN  
L2 1160 S 3068.74/RID  
L3 451970 S 46.383/RID  
L4 226 S L2 AND L3

FILE 'CAPLUS' ENTERED AT 15:56:26 ON 11 JUN 2003

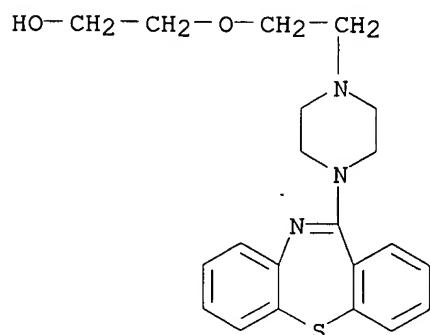
L5 452 S L4  
L6 178 S L1  
L7 1388054 S WEIGHT  
L8 27 S L5 AND L7  
L9 19 S L6 AND L7  
L10 27 S L8 OR L9  
L11 23143 S OBESITY  
L12 6 S L5 AND L11  
L13 6 S L6 AND L11  
L14 30 S L10 OR L12 OR L13  
L15 79170 S DIABETES  
L16 4021 S PSYCHOSIS  
L17 7 S L15 AND L5  
L18 5 S L15 AND L6  
L19 38 S L16 AND L5  
L20 30 S L16 AND L6  
L21 7 S L17 OR L18  
L22 38 S L19 OR L20  
L23 44 S L21 OR L22  
L24 66 S L14 OR L23  
L25 26 S L24 AND PATENT/DT  
L26 40 S L24 NOT L25  
L27 0 S L26 AND 2003/SO  
L28 14 S L26 AND 2002/SO  
L29 9 S L26 AND 2001/SO  
L30 43 S L24 NOT (L28 OR L29)

=> d scan l1

YOU HAVE REQUESTED DATA FROM FILE 'REGISTRY' - CONTINUE? (Y)/N:y

10/009,574

L1 1 ANSWERS REGISTRY COPYRIGHT 2003 ACS  
IN Ethanol, 2-[2-(4-dibenzo[b,f][1,4]thiazepin-11-yl-1-piperazinyl)ethoxy]-  
(9CI)  
MF C21 H25 N3 O2 S  
CI COM



\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

ALL ANSWERS HAVE BEEN SCANNED

10/009,574

=> d bib abs hitstr 1-43 130

L30 ANSWER 1 OF 43 CAPLUS COPYRIGHT 2003 ACS

AN 2003:396456 CAPLUS

TI Substance to prevent or reverse **weight** gain induced by psychoactive agents

IN Miller, Jon M.

PA USA

SO U.S. Pat. Appl. Publ., 5 pp.

CODEN: USXXCO

DT Patent

LA English

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	US 2003096808	A1	20030522	US 1999-280279	19990329
PRAI	US 1999-280279		19990329		

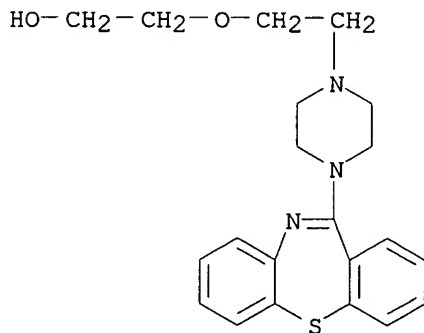
AB A substance to prevent or reverse **wt.** gain induced by psychoactive agents comprises an antipsychotic drug or mood stabilizing drug in a concn. from 0.01% to 99.99% in combination with a histamine H2-receptor antagonist in a concn. from 99.99% to 0.01%. Example antipsychotic drugs are olanzapine, clozapine, risperidone, and quetiapine. The antipsychotic drug is typically in a concn. of 10% to 90%, 30% to 60% and 50%. Example mood stabilizing drugs are divalproex sodium, valproic acid, and mirtazapine. The mood stabilizing drug is typically in a concn. of 10% to 90%, 30% to 60% and 50%. Example histamine H2-receptor antagonist are nizatidine, famotidine, cimetidine and ranitidine. The histamine H2-receptor antagonist (16) is typically in a concn. of 60% to 30% and 50%.

IT **111974-69-7**, Quetiapine

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (antipsychotic; substance to prevent or reverse **wt.** gain induced by psychoactive agents)

RN 111974-69-7 CAPLUS

CN Ethanol, 2-[2-(4-dibenzo[b,f][1,4]thiazepin-11-yl-1-piperazinyl)ethoxy]-  
 (9CI) (CA INDEX NAME)



L30 ANSWER 2 OF 43 CAPLUS COPYRIGHT 2003 ACS

AN 2003:319255 CAPLUS

DN 138:343854

TI Buccal sprays or capsules containing drugs for treating disorders of the central nervous system

IN Dugger, Harry A.

PA USA

SO U.S. Pat. Appl. Publ., 17 pp., Cont.-in-part of U.S. Ser. No. 537,118.  
CODEN: USXXCO

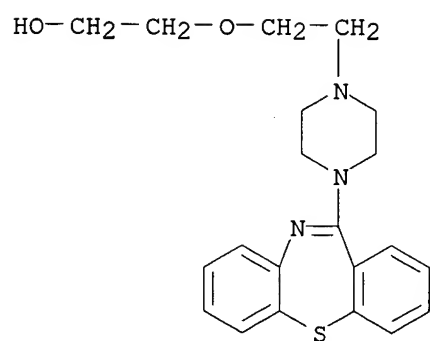
DT Patent

LA English

FAN.CNT 8

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	US 2003077227	A1	20030424	US 2002-230060	20020829
	WO 9916417	A1	19990408	WO 1997-US17899	19971001
	W:		AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GE, GH, HU, IL, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM		
	RW:		GH, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG		
	EP 1029536	A1	20000823	EP 2000-109347	19971001
	R:		AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO		
	EP 1036561	A1	20000920	EP 2000-109357	19971001
	R:		AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO		
PRAI	WO 1997-US17899	A2	19971001		
	US 2000-537118	A2	20000329		
	EP 1997-911621	A3	19971001		
AB	Buccal aerosol sprays or capsules using polar and non-polar solvent have now been developed which provide biol. active compds. for rapid absorption through the oral mucosa, resulting in fast onset of effect. The buccal polar compns. of the invention comprise formulation A: aq. polar solvent, active compd., and optional flavoring agent; formulation B: aq. polar solvent, active compd., optionally flavoring agent, and propellant; formulation C: non-polar solvent, active compd., and optional flavoring agent; and formulation D: non-polar solvent, active compd., optional flavoring agent, and propellant. Thus, a lingual spray contained sumatriptan succinate 10-15, EtOH 10-20, propylene glycol 10-15, PEG 35-40, water 10-15, and flavors 2-3%.				
IT	<b>111974-69-7, Quetiapine</b>				
	RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (buccal sprays or capsule contg. drugs for treating disorders of central nervous system)				
RN	111974-69-7 CAPLUS				
CN	Ethanol, 2-[2-(4-dibenzo[b,f][1,4]thiazepin-11-yl-1-piperazinyl)ethoxy]-(9CI) (CA INDEX NAME)				

10/009,574



L30 ANSWER 3 OF 43 CAPLUS COPYRIGHT 2003 ACS

AN 2003:261599 CAPLUS

DN 138:265698

TI Organic acid-conjugated antipsychotic drugs, and therapeutic use thereof

IN Nudelman, Abraham; Rephaeli, Ada; Gil-Ad, Irit; Weizman, Abraham

PA Ramot at Tel Aviv University Ltd., Israel; Bar Ilan University

SO PCT Int. Appl., 107 pp.

CODEN: PIXXD2

DT Patent

LA English

FAN.CNT 1

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2003026563	A2	20030403	WO 2002-IL795	20020929
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW, AM, AZ, BY RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				

PRAI US 2001-324936P P 20010927

AB Chem. conjugates of anti-psychotic drugs and org. acids, uses thereof in the treatment of psychotic and/or proliferative disorders and diseases and as chemosensitizing agents, and their syntheses, are disclosed. The org. acids are selected to reduce side effects induced by the anti-psychotic drugs and/or to exert an anti-proliferative activity.

IT 2058-52-8D, Clothiapine, org. acid conjugates 111974-69-7D

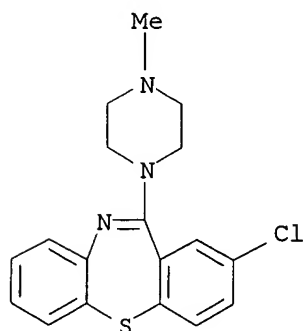
, Quetiapine, org. acid conjugates

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(org. acid-conjugated antipsychotic drugs, and therapeutic use)

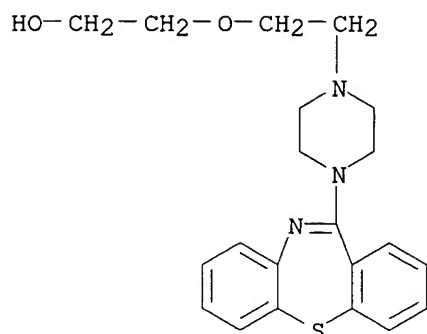
RN 2058-52-8 CAPLUS

CN Dibenzo[b,f][1,4]thiazepine, 2-chloro-11-(4-methyl-1-piperazinyl)- (7CI, 8CI, 9CI) (CA INDEX NAME)

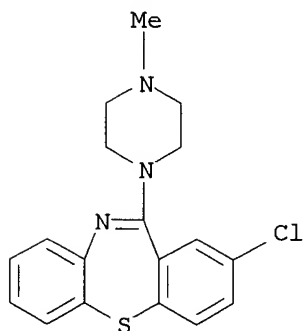


RN 111974-69-7 CAPLUS

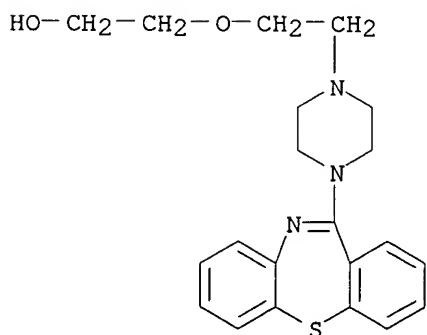
CN Ethanol, 2-[2-(4-dibenzo[b,f][1,4]thiazepin-11-yl-1-piperazinyl)ethoxy]- (9CI) (CA INDEX NAME)



IT 2058-52-8, Clothiapine 111974-69-7, Quetiapine  
 RL: RCT (Reactant); RACT (Reactant or reagent)  
 (org. acid-conjugated antipsychotic drugs, and therapeutic use)  
 RN 2058-52-8 CAPLUS  
 CN Dibenzo[b,f][1,4]thiazepine, 2-chloro-11-(4-methyl-1-piperazinyl)- (7CI, 8CI, 9CI) (CA INDEX NAME)



RN 111974-69-7 CAPLUS  
 CN Ethanol, 2-[2-(4-dibenzo[b,f][1,4]thiazepin-11-yl-1-piperazinyl)ethoxy]- (9CI) (CA INDEX NAME)



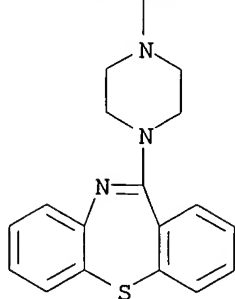
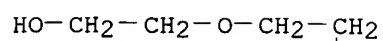


10/009,574

L30 ANSWER 4 OF 43 CAPLUS COPYRIGHT 2003 ACS  
 AN 2003:97307 CAPLUS  
 DN 138:147748  
 TI Methods for preventing antipsychotic-induced **weight** gain  
 IN Belanoff, Joseph K.; Schatzberg, Alan F.  
 PA Corcept Therapeutics, Inc., USA  
 SO PCT Int. Appl., 32 pp.  
 CODEN: PIXXD2  
 DT Patent  
 LA English  
 FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 2003009853	A1	20030206	WO 2002-US23441	20020722
	W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				
	US 2003027802	A1	20030206	US 2002-201356	20020722
PRAI	US 2001-307693P	P	20010723		
AB	This invention generally pertains to the field of psychiatry. In particular, this invention pertains to the discovery that agents capable of inhibiting the binding of cortisol to its receptors can be used in methods for preventing antipsychotic-induced <b>wt.</b> gain. Mifepristone, a potent specific glucocorticoid receptor antagonist, can be used in these methods. The invention also provides a kit for preventing antipsychotic-induced <b>wt.</b> gain in a human including a glucocorticoid receptor antagonist and instructional material teaching the indications, dosage and schedule of administration of the glucocorticoid receptor antagonist.				
IT	<b>111974-69-7</b> , Quetiapine RL: ADV (Adverse effect, including toxicity); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (glucocorticoid receptor antagonist for prevention and reversal of antipsychotic-induced <b>wt.</b> gain)				
RN	111974-69-7 CAPLUS				
CN	Ethanol, 2-[2-(4-dibenzo[b,f][1,4]thiazepin-11-yl-1-piperazinyl)ethoxy]-(9CI) (CA INDEX NAME)				

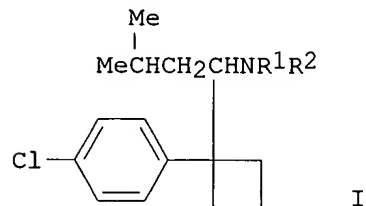
10/009,574



RE.CNT 1      THERE ARE 1 CITED REFERENCES AVAILABLE FOR THIS RECORD  
ALL CITATIONS AVAILABLE IN THE RE FORMAT

L30 ANSWER 5 OF 43 CAPLUS COPYRIGHT 2003 ACS  
 AN 2003:23534 CAPLUS  
 DN 138:66716  
 TI Method of controlling **weight** gain associated with therapeutic drugs  
 IN Mendel, Carl M.; Seaton, Timothy B.; Weinstein, Steve P.  
 PA USA  
 SO U.S. Pat. Appl. Publ., 5 pp.  
 CODEN: USXXCO  
 DT Patent  
 LA English  
 FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	US 2003008897	A1	20030109	US 2000-527813	20000317
PRAI	US 2000-527813		20000317		
OS	MARPAT 138:66716				
GI					



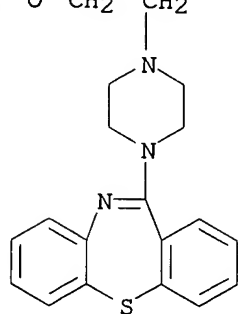
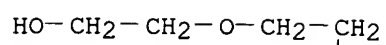
AB The invention discloses the use of compd. I [R<sup>1</sup>,R<sup>2</sup> = H or Methyl] for treating **wt.** gain assocd. with treatment with certain drugs including the tricyclic antidepressants, lithium, sulfonylureas, beta-adrenergic blockers, certain steroid contraceptives, corticosteroids, insulin, cyproheptadine, sodium valproate, neuroleptics, phenothiazine or pizotifen.

IT **111974-69-7**, Quetiapine  
 RL: ADV (Adverse effect, including toxicity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (method of controlling **wt.** gain assocd. with therapeutic drugs)

RN 111974-69-7 CAPLUS

CN Ethanol, 2-[2-(4-dibenzo[b,f][1,4]thiazepin-11-yl-1-piperazinyl)ethoxy]-(9CI) (CA INDEX NAME)

10/009,574



~~D30~~ ANSWER 6 OF 43 CAPLUS COPYRIGHT 2003 ACS

~~AN~~ 2003:5941 CAPLUS

~~DN~~ 138:73274

TI Preparation of substituted piperazinyldibenzo[b,f][1,4]oxazepines and thiazepines as atypical antipsychotic agents having low affinity for the D2-receptor

IN Kapur, Shitij; McClelland, Robert

PA Neuromolecular, Inc., Can.

SO PCT Int. Appl., 76 pp.

CODEN: PIXXD2

DT Patent

LA English

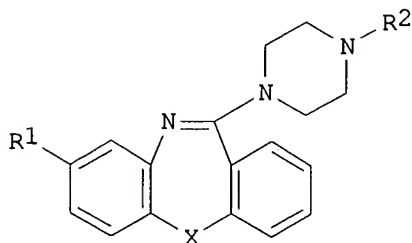
FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 2003000670	A1	20030103	WO 2002-CA956	20020626
	W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				

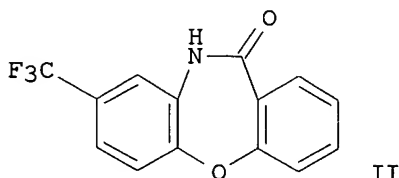
PRAI US 2001-300430P P 20010626

OS MARPAT 138:73274

GI



I



II

AB Title compds. I [R1 = halo, CF3, CF3O, CN, CH3, CH3O; R2 = alkyl, etc.; X = O, S] are prepd. For instance, Me salicylate was reacted with 4-fluoro-3-nitrobenzotrifluoride (CH3CN, 18-crown-6, 40% wt./wt. KF-alumina, reflux, 4 h) to afford Me 2-((2-nitro-4-trifluoromethylphenyl)oxy)benzoate. This intermediate was reduced to the amino deriv., sapond. and cyclized (xylene, reflux, 24 h) to II. II was treated with POCl3 to afford the imino chloride intermediate and subsequently treated with 1-ethylpiperazine to afford I [R1 = CF3; R2 = Et; X = O; III]. III had Ki = 258 nM for the D2 receptor. I are useful for the treatment of psychiatric disorders (e.g., **psychosis**, depression, schizophrenia).

IT 479681-02-2P 479681-11-3P 479681-16-8P  
479681-19-1P 479681-23-7P 479681-26-0P  
479681-30-6P 479681-57-7P

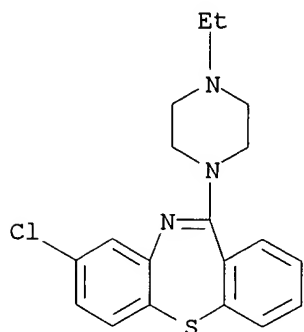
RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU

(Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(prepn. of substituted piperazinyldibenzo[b,f][1,4]oxazepines and thiazepines as atypical antipsychotic agents having low affinity for D2-receptor)

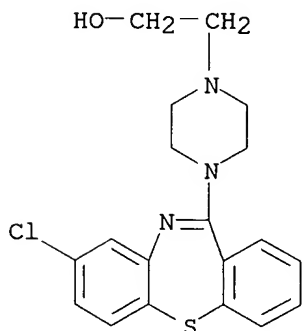
RN 479681-02-2 CAPLUS

CN Dibenzo[b,f][1,4]thiazepine, 8-chloro-11-(4-ethyl-1-piperazinyl)- (9CI)  
(CA INDEX NAME)



RN 479681-11-3 CAPLUS

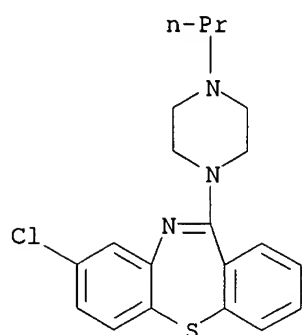
CN 1-Piperazineethanol, 4-(8-chlorodibenzo[b,f][1,4]thiazepin-11-yl)- (9CI)  
(CA INDEX NAME)



RN 479681-16-8 CAPLUS

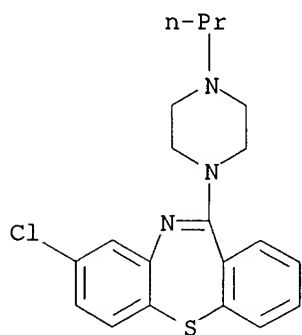
CN Dibenzo[b,f][1,4]thiazepine, 8-chloro-11-(4-propyl-1-piperazinyl)-, monohydrochloride (9CI) (CA INDEX NAME)

10/009,574



● HCl

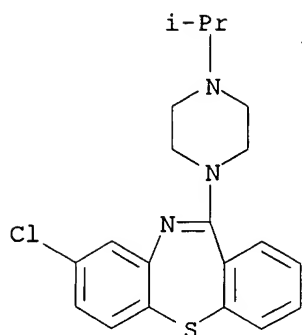
RN 479681-19-1 CAPLUS  
CN Dibenzo[b,f][1,4]thiazepine, 8-chloro-11-(4-propyl-1-piperazinyl)- (9CI)  
(CA INDEX NAME)



RN 479681-23-7 CAPLUS  
CN Dibenzo[b,f][1,4]thiazepine, 8-chloro-11-[4-(1-methylethyl)-1-piperazinyl]-  
, monohydrochloride (9CI) (CA INDEX NAME)

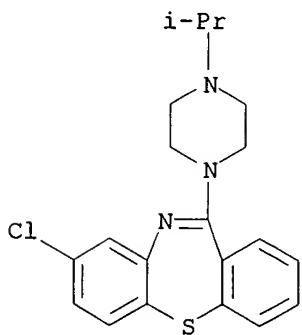


10/009,574



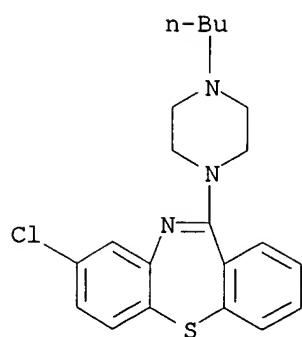
● HCl

RN 479681-26-0 CAPLUS  
CN Dibenzo[b,f][1,4]thiazepine, 8-chloro-11-[4-(1-methylethyl)-1-piperazinyl]-  
(9CI) (CA INDEX NAME)



RN 479681-30-6 CAPLUS  
CN Dibenzo[b,f][1,4]thiazepine, 11-(4-butyl-1-piperazinyl)-8-chloro-,  
monohydrochloride (9CI) (CA INDEX NAME)

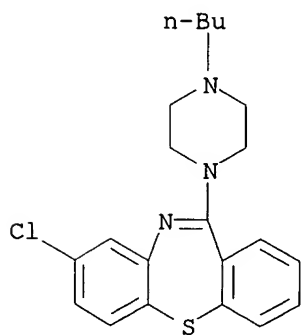
10/009,574



● HCl

RN 479681-57-7 CAPLUS

CN Dibenzo[b,f][1,4]thiazepine, 11-(4-butyl-1-piperazinyl)-8-chloro- (9CI)  
(CA INDEX NAME)



RE.CNT 7 THERE ARE 7 CITED REFERENCES AVAILABLE FOR THIS RECORD  
ALL CITATIONS AVAILABLE IN THE RE FORMAT

L30 ANSWER 7 OF 43 CAPLUS COPYRIGHT 2003 ACS

AN 2002:977588 CAPLUS

DN 138:33362

TI Use of cyclooxygenase 2 (COX-2) inhibitors for the treatment of schizophrenia, delusional disorders, affective disorders, autism, or tic disorders

IN Muller, Norbert

PA Germany

SO PCT Int. Appl., 58 pp.

CODEN: PIXXD2

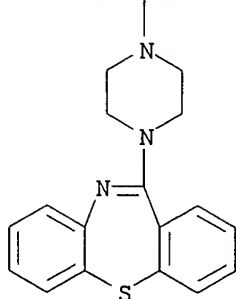
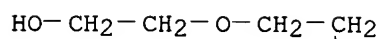
DT Patent

LA English

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 2002102297	A2	20021227	WO 2002-EP6013	20020531
	WO 2002102297	A3	20030501		
	W:	AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM			
	RW:	GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG			
	DE 10129320	A1	20030410	DE 2001-10129320	20010619
PRAI	DE 2001-10129320	A	20010619		
	US 2002-364904P	P	20020314		
AB	The invention discloses the use of a COX-2 inhibitor for the treatment of psychiatric disorders, e.g. schizophrenia, delusional disorders, affective disorders, autism or tic disorders, in particular chronic schizophrenic <b>psychoses</b> and schizoaffective <b>psychoses</b> , temporary acute psychotic disorders, depressive episodes, recurring depressive episodes, manic episodes and bipolar affective disorders. Moreover, the invention discloses the use of a COX-2 inhibitor, in particular celecoxib, in combination with a neuroleptic drug, in particular risperidone, or an antidepressant, for the treatment of psychiatric disorders such as schizophrenia, delusional disorders, affective disorders, autism or tic disorders.				
IT	<b>111974-69-7</b> , Quetiapine <b>111974-72-2</b> , Quetiapine fumarate RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (cyclooxygenase 2 inhibitors for treatment of psychiatric disorders, and use with other agents)				
RN	111974-69-7 CAPLUS				
CN	Ethanol, 2-[2-(4-dibenzo[b,f][1,4]thiazepin-11-yl-1-piperazinyl)ethoxy]-(9CI) (CA INDEX NAME)				

10/009,574



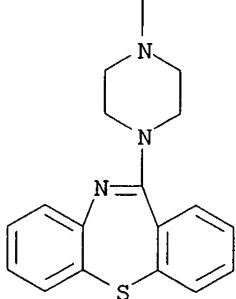
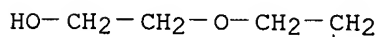
RN 111974-72-2 CAPLUS

CN Ethanol, 2-[2-(4-dibenzo[b,f][1,4]thiazepin-11-yl-1-piperazinyl)ethoxy]-, (2E)-2-butenedioate (2:1) (salt) (9CI) (CA INDEX NAME)

CM 1

CRN 111974-69-7

CMF C21 H25 N3 O2 S

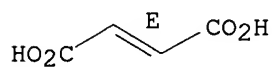


CM 2

CRN 110-17-8

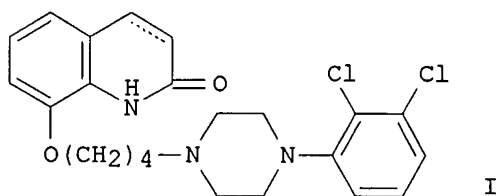
CMF C4 H4 O4

Double bond geometry as shown.



L30 ANSWER 8 OF 43 CAPLUS COPYRIGHT 2003 ACS  
 AN 2002:889556 CAPLUS  
 DN 137:363096  
 TI Carbostyryl derivative 5-HT1a receptor subtype agonist for treatment of  
 central nervous system disorders  
 IN Jordan, Shaun; Kikuchi, Tetsuro; Tottori, Katsura; Hirose, Tsuyoshi;  
 Uwahodo, Yasufumi  
 PA USA  
 SO U.S. Pat. Appl. Publ., 8 pp.  
 CODEN: USXXCO  
 DT Patent  
 LA English  
 FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	US 2002173513	A1	20021121	US 2002-55915	20020128
PRAI	US 2001-331370P	P	20010129		
GI					

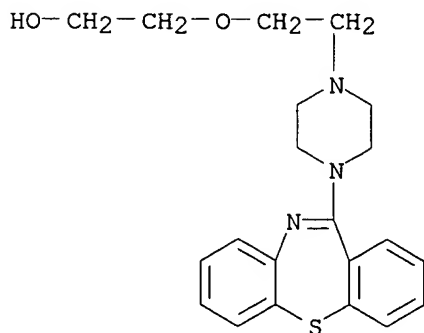


AB The invention provides a method for treating a patient suffering from a disorder of the central nervous system assocd. with the 5-HT1a receptor subtype, comprising as an active ingredient a carbostyryl deriv. I (carbon-carbon bond between 3- and 4-positions in carbostyryl skeleton is single or double bond), or a salt thereof.

IT **111974-69-7, Quetiapine**  
 RL: BSU (Biological study, unclassified); BIOL (Biological study)  
 (carbostyryl deriv. 5-HT1a receptor subtype agonist for treatment of central nervous system disorders)

RN 111974-69-7 CAPLUS

CN Ethanol, 2-[2-(4-dibenzo[b,f][1,4]thiazepin-11-yl-1-piperazinyl)ethoxy]-  
 (9CI) (CA INDEX NAME)



10/009,574

L30 ANSWER 9 OF 43 CAPLUS COPYRIGHT 2003 ACS

AN 2002:849447 CAPLUS

DN 137:333167

TI Treatment of psychotic disorders using co-therapy with anticonvulsant derivatives and atypical antipsychotics

IN Fenton, Wayne S.

PA Ortho-McNeil Pharmaceutical, Inc., USA

SO PCT Int. Appl., 26 pp.

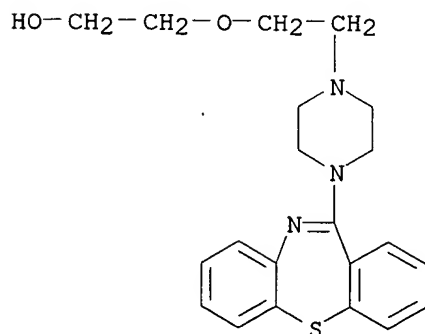
CODEN: PIXXD2

DT Patent

LA English

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 2002087590	A1	20021107	WO 2002-US12997	20020423
	W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, UZ, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				
PRAI	US 2001-286765P	P	20010426		
	US 2001-301661P	P	20010628		
OS	MARPAT 137:333167				
AB	Treatment of psychotic disorders (e.g. schizophrenia; schizophreniform and schizoaffective disorders) comprises co-therapy with an anticonvulsant deriv. (e.g. topiramate) and atypical antipsychotic (e.g. olanzapine).				
IT	111974-69-7, Quetiapine				
	RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)				
	(anticonvulsant deriv.-atypical antipsychotic co-therapy for psychotic disorders)				
RN	111974-69-7 CAPLUS				
CN	Ethanol, 2-[2-(4-dibenzo[b,f][1,4]thiazepin-11-yl-1-piperazinyl)ethoxy]-(9CI) (CA INDEX NAME)				



RE.CNT 11 THERE ARE 11 CITED REFERENCES AVAILABLE FOR THIS RECORD  
ALL CITATIONS AVAILABLE IN THE RE FORMAT

L30 ANSWER 10 OF 43 CAPLUS COPYRIGHT 2003 ACS

AN 2002:754219 CAPLUS

DN 137:273219

TI Anti-**psychosis** combination containing a modulator of 5-HT2A receptor

IN Behan, Dominic P.; Chalmers, Derek T.; Menzaghi, Frederique

PA Arena Pharmaceuticals, Inc., USA

SO PCT Int. Appl., 54 pp.

CODEN: PIXXD2

DT Patent

LA English

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 2002076464	A1	20021003	WO 2002-US9086	20020322
	W:	AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM			
	RW:	GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG			
	US 2002156068	A1	20021024	US 2002-104602	20020322
PRAI	US 2001-278516P	P	20010322		

OS MARPAT 137:273219

AB This invention relates to methods of reducing hyperlocomotor activity and stereotypy by administering a compn. comprising a modulator of the 5-HT2A receptor with a neuroleptic agent used for treating **psychoses**, such as Haloperidol. The invention further relates to compns. comprising a modulator of the 5-HT2A receptor with a neuroleptic agent. For example, a 5-HT2A receptor modulator N-[3-(4-bromo-2-methylpyrazol-3-yl)phenyl] [(4-chlorophenyl)amino]carboxamide (AR 116081) potentiated the effect of the neuroleptic haloperidol in a model of **psychosis** in rats. Thus, in combination, modulators of the 5-HT2A receptor, preferably AR116081, and neuroleptics, preferably haloperidol, preferably at a low dosage, will reverse the hyperactivity in the rat model, thereby potentially reducing the side effects usually assocd. with neuroleptics (e.g., extrapyramidal motor syndrome and tardive dyskinesia).

IT 111974-69-7, Quetiapine

RL: ADV (Adverse effect, including toxicity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

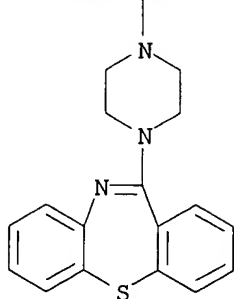
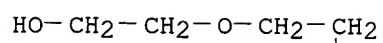
(anti-**psychosis** combination contg. modulator of 5-HT2A receptor)

RN 111974-69-7 CAPLUS

CN Ethanol, 2-[2-(4-dibenzo[b,f][1,4]thiazepin-11-yl-1-piperazinyl)ethoxy]-(9CI) (CA INDEX NAME)



10/009,574



RE.CNT 1      THERE ARE 1 CITED REFERENCES AVAILABLE FOR THIS RECORD  
ALL CITATIONS AVAILABLE IN THE RE FORMAT

L30 ANSWER 11 OF 43 CAPLUS COPYRIGHT 2003 ACS

AN 2002:674788 CAPLUS

DN 137:195595

TI Atypical antipsychotic-antidepressant combination for treatment of depression, obsessive compulsive disorder, and **psychosis**

IN Howard, Harry R., Jr.

PA Pfizer Inc., USA

SO U.S. Pat. Appl. Publ., 20 pp.

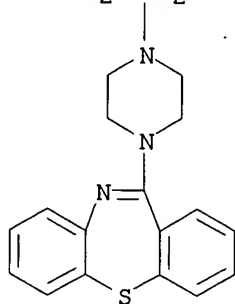
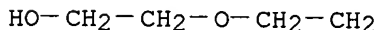
CODEN: USXXCO

DT Patent

LA English

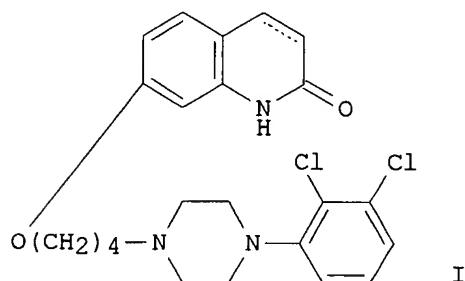
FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	US 2002123490	A1	20020905	US 2001-10651	20011206
	EP 1238676	A1	20020911	EP 2002-251153	20020220
	R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR				
	JP 2002308801	A2	20021023	JP 2002-50579	20020227
PRAI	US 2001-272619P	P	20010301		
OS	MARPAT 137:195595				
AB	The invention provides a method for treating depression, obsessive compulsive disorder, and <b>psychosis</b> in a mammal, including a human, by administering to the mammal an atypical antipsychotic in combination with an antidepressant agent with improvement in efficiency. It also provides pharmaceutical compns. contg. a pharmaceutically acceptable carrier, an atypical antipsychotic, and a serotonin reuptake inhibitor.				
IT	<b>111974-69-7</b> , Quetiapine				
	RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)				
	(atypical antipsychotic-antidepressant combination for treatment of depression, obsessive compulsive disorder, and <b>psychosis</b> )				
RN	111974-69-7 CAPLUS				
CN	Ethanol, 2-[2-(4-dibenzo[b,f][1,4]thiazepin-11-yl-1-piperazinyl)ethoxy]-(9CI) (CA INDEX NAME)				



L30 ANSWER 12 OF 43 CAPLUS COPYRIGHT 2003 ACS  
 AN 2002:594663 CAPLUS  
 DN 137:150248  
 TI Carbostyryl derivative 5-HT1a receptor agonists for treatment of central nervous system disorders  
 IN Jordan, Shaun; Kikuchi, Tetsuro; Tottori, Katsura; Hirose, Tsuyoshi; Uwahodo, Yasufumi  
 PA Otsuka Pharmaceutical Co., Ltd., Japan  
 SO PCT Int. Appl., 31 pp.  
 CODEN: PIXXD2  
 DT Patent  
 LA English  
 FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 2002060423	A2	20020808	WO 2002-JP626	20020129
	WO 2002060423	A3	20030410		
	W: AU, BR, CA, CN, ID, IN, JP, KR, MX, PH, SG				
	RW: AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR				
PRAI	US 2001-770210	A	20010129		
GI					



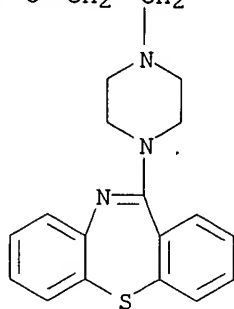
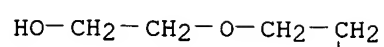
AB The invention discloses the use of a compd. for the prodn. of a medicament for treating a patient suffering from a disorder of the central nervous system assocd. with 5-HT1a receptor subtype, the medicament including as an active ingredient a carbostyryl deriv. I (C-C bond between 3- and 4-positions in the carbostyryl skeleton is single or double bond), or a pharmaceutically acceptable salt or solvate thereof.

IT **111974-69-7**, Quetiapine  
 RL: BSU (Biological study, unclassified); PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (carbostyryl deriv. 5-HT1a receptor agonists for treatment of central nervous system disorders)

RN 111974-69-7 CAPLUS

CN Ethanol, 2-[2-(4-dibenzo[b,f][1,4]thiazepin-11-yl-1-piperazinyl)ethoxy]-(9CI) (CA INDEX NAME)

10/009,574



L30 ANSWER 13 OF 43 CAPLUS COPYRIGHT 2003 ACS

AN 2002:521465 CAPLUS

DN 137:98994

TI Pharmaceuticals containing a combination of norepinephrine reuptake inhibitors and neuroleptics

IN Wong, Erik Ho Fong; Gallen, Christopher C.; Svensson, Torgny

PA Pharmacia &amp; Upjohn Company, USA; Pharmacia AB

SO PCT Int. Appl., 22 pp.

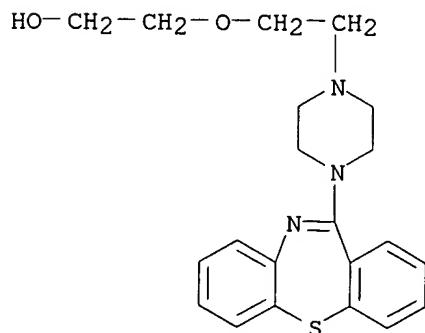
CODEN: PIXXD2

DT Patent

LA English

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 2002053140	A2	20020711	WO 2001-US45871	20011227
	WO 2002053140	A3	20021024		
	W:	AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM			
	RW:	GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG			
	US 2002156067	A1	20021024	US 2001-35100	20011228
PRAI	US 2001-259286P	P	20010102		
AB	A compn. comprising: (a) a pharmaceutically effective amt. of one or more norepinephrine reuptake inhibitors or a salt; and (b) 1 or more neuroleptics is provided. The compn. is useful in treating disorders or diseases of the central nervous system, and particularly useful in treating schizophrenia. A pharmaceutical compn. was prepd. by combining reboxetine with a neuroleptic in an acceptable carrier. The compn. contains 0.01-10 mg rebexetine and 25-300 mg clozapine.				
IT	111974-69-7, Quetiapine				
	RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)				
	(pharmaceuticals contg. combination of norepinephrine reuptake inhibitors and neuroleptics)				
RN	111974-69-7 CAPLUS				
CN	Ethanol, 2-[2-(4-dibenzo[b,f][1,4]thiazepin-11-yl-1-piperazinyl)ethoxy]-(9CI) (CA INDEX NAME)				



10/009,574

L30 ANSWER 14 OF 43 CAPLUS COPYRIGHT 2003 ACS

AN 2002:240731 CAPLUS

DN 136:257287

TI Compounds and methods for diagnosing and treating amyloid-related conditions

IN Raub, Thomas J.; Tanis, Steven P.; Buhl, Allen Edwin; Carter, Donald Bainbridge; Bandiera, Tiziano; Lansen, Jacqueline; Pellerano, Cesare; Savini, Luisa

PA Pharmacia &amp; Upjohn Company, USA; Pharmacia &amp; Upjohn S.p.A.

SO PCT Int. Appl., 56 pp.

CODEN: PIXXD2

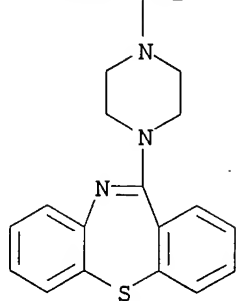
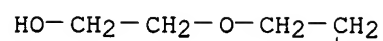
DT Patent

LA English

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 2002024652	A1	20020328	WO 2001-US29010	20010917
	WO 2002024652	B1	20020627		
	W:	AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM			
	RW:	GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG			
	AU 2001089123	A5	20020402	AU 2001-89123	20010917
PRAI	US 2000-234611P	P	20000922		
	US 2000-667357	A	20000922		
	WO 2001-US29010	W	20010917		
OS	MARPAT 136:257287				
AB	The invention provides methods for diagnosing and treating amyloid-related conditions and compds. useful for the same. The invention provides for detecting, imaging, monitoring, diagnosing, and treating conditions characterized by the binding or aggregation of amyloid fibrils. More particularly, the invention relates to using quinolinehydrazones compds. for diagnosing and treating amyloidotic conditions and also as an antioxidant. Examples are provided showing that 4-methyl-7-methoxy-2-(4-quinolylmethylenehydrazino)quinoline is suitable for fluorescence detection of amyloid plaque and has antioxidant activity.				
IT	<b>111974-69-7</b> , Quetiapine				
	RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (quinolinehydrazones compds. for diagnosing and assessing treatment of amyloidotic conditions)				
RN	111974-69-7 CAPLUS				
CN	Ethanol, 2-[2-(4-dibenzo[b,f][1,4]thiazepin-11-yl-1-piperazinyl)ethoxy]-(9CI) (CA INDEX NAME)				

10/009,574



RE.CNT 7

THERE ARE 7 CITED REFERENCES AVAILABLE FOR THIS RECORD  
ALL CITATIONS AVAILABLE IN THE RE FORMAT



L30 ANSWER 15 OF 43 CAPLUS COPYRIGHT 2003 ACS  
 AN 2002:153684 CAPLUS  
 DN 136:194261  
 TI Therapeutic combinations of (S)-2-(benzylamino-methyl)-2,3,8,9,-tetrahydro  
 7H-1,4-dioxino{2,3-e}indol-8-one and neuroleptics for the treatment or  
 prevention of psychotic disorders  
 IN Marquis, Karen L.  
 PA American Home Products Corporation, USA  
 SO U.S., 8 pp.  
 CODEN: USXXAM  
 DT Patent  
 LA English  
 FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	US 6350773	B1	20020226	US 2000-728994	20001204
PRAI	US 1999-240908P	P	19991210		

AB Therapeutic combinations useful in the treatment or prevention of  
 psychotic disorders, to pharmaceutical compns. contg. said combinations,  
 and to their use in the treatment or prophylaxis of prevention disorders  
 are provided. The effect of (S)-2-(benzylamino-methyl)-2,3,8,9-tetrahydro-  
 7H-1,4-dioxino[2,3-e]indol-8-one on haloperidol-induced catalepsy in rats  
 at 60 min after drug treatment was studied. A dose-dependent decrease in  
 time spent in catalepsy position was obsd. A minimal ED of 0.3 mg/kg and  
 an ED50 (dose producing 50% redn. in maximal response) of 0.08 mg/kg were  
 calcd. from these results.

IT **111974-72-2, Seroquel**  
 RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL  
 (Biological study); USES (Uses)  
 (therapeutic combinations of benzylaminotetrahydrodioxinoindolone and  
 neuroleptics for treatment or prevention of psychotic disorders)

RN 111974-72-2 CAPLUS

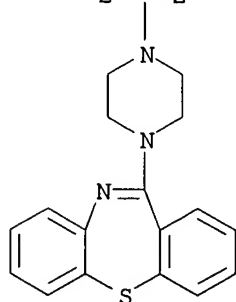
CN Ethanol, 2-[2-(4-dibenzo[b,f][1,4]thiazepin-11-yl-1-piperazinyl)ethoxy]-,  
 (2E)-2-butenedioate (2:1) (salt) (9CI) (CA INDEX NAME)

CM 1

CRN 111974-69-7

CMF C21 H25 N3 O2 S

HO-CH<sub>2</sub>-CH<sub>2</sub>-O-CH<sub>2</sub>-CH<sub>2</sub>

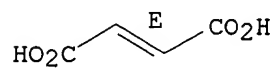


CM 2

10/009,574

CRN 110-17-8  
CMF C4 H4 O4

Double bond geometry as shown.



RE.CNT 8      THERE ARE 8 CITED REFERENCES AVAILABLE FOR THIS RECORD  
ALL CITATIONS AVAILABLE IN THE RE FORMAT

L30 ANSWER 16 OF 43 CAPLUS COPYRIGHT 2003 ACS

AN 2001:525912 CAPLUS

DN 135:112000

TI Osmotic device containing venlafaxine and an anti-psychotic agent

IN Faour, Joaquina; Vergez, Juan A.

PA Laboratorios Phoenix U.S.A., Inc., USA

SO PCT Int. Appl., 39 pp.

CODEN: PIXXD2

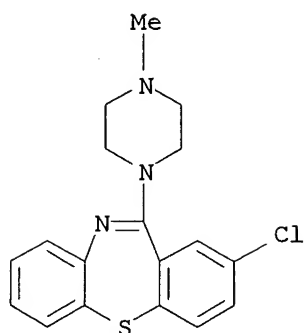
DT Patent

LA English

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 2001051041	A1	20010719	WO 2001-US580	20010108
	W:				
					AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
	RW:				GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
	US 2001048943	A1	20011206	US 2000-728276	20001130
	US 6572890	B2	20030603		
	EP 1246614	A1	20021009	EP 2001-901877	20010108
	R:				AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR
PRAI	US 2000-175822P	P	20000113		
	US 2000-728276	A	20001130		
	WO 2001-US580	W	20010108		
AB	The present invention provides an osmotic device contg. controlled release venlafaxine in the core in combination with an anti-psychotic agent in a rapid release external coat. A wide range of anti-psychotic agents can be used in this device. Particular embodiments of the invention provide osmotic devices having predetd. release profiles. One embodiment of the osmotic device includes an external coat that has been spray-coated rather than compression-coated onto the device. The device with spray-coated external core is smaller and easier to swallow than the similar device having a compression-coated external coat. The device is useful for the treatment of depression anxiety or <b>psychosis</b> related disorders. Thus, a core formulation contained venlafaxine 10-500, osmagent 17-250, binder 7.5-50, plasticizer (low mol. wt.) 0.1-25, glidant 0.1-6, plasticizer (high mol. wt.) 2.5-30, and lubricant 1-7.5 mg. Water sol. polymers were used in the coating formulations.				
IT	<b>2058-52-8</b> , Clothiapine <b>111974-69-7</b> , Quetiapine				
	RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (osmotic device contg. venlafaxine and anti-psychotic agent)				
RN	2058-52-8 CAPLUS				
CN	Dibenzo[b,f][1,4]thiazepine, 2-chloro-11-(4-methyl-1-piperazinyl)- (7CI, 8CI, 9CI) (CA INDEX NAME)				

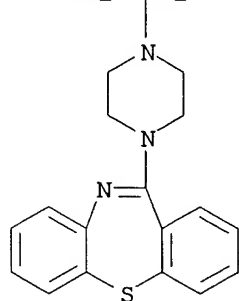
10/009,574



RN 111974-69-7 CAPLUS

CN Ethanol, 2-[2-(4-dibenzo[b,f][1,4]thiazepin-11-yl-1-piperazinyl)ethoxy]-  
(9CI) (CA INDEX NAME)

HO-CH<sub>2</sub>-CH<sub>2</sub>-O-CH<sub>2</sub>-CH<sub>2</sub>



RE.CNT 3 THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD  
ALL CITATIONS AVAILABLE IN THE RE FORMAT

L30 ANSWER 17 OF 43 CAPLUS COPYRIGHT 2003 ACS

AN 2001:525911 CAPLUS

DN 135:111999

TI Osmotic device containing alprazolam and an antipsychotic agent

IN Faour, Joaquina; Vergez, Juan A.

PA Laboratorios Phoenix U.S.A., Inc., USA

SO PCT Int. Appl., 38 pp.

CODEN: PIXXD2

DT Patent

LA English

FAN.CNT 1

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2001051040	A1	20010719	WO 2001-US637	20010109
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				
US 2002051807	A1	20020502	US 2001-756497	20010108

PRAI US 2000-175827P P 20000113

AB The present invention provides an osmotic device contg. controlled release alprazolam in the core optionally in combination with an anti-psychotic agent, in a rapid release external coat. A wide range of anti-psychotic agents can be used in this device. Particular embodiments of the invention provide osmotic devices having predetd. release profiles. One preferred embodiment of the osmotic device includes an external coat that has been spray coated rather than compression coated onto the device. The device with spray coated external coat is smaller and easier to swallow than the similar device having a compression coated external coat. The device is useful for the treatment of depression, anxiety or **psychosis** related disorders. Thus, osmotic-release tablets contained alprazolam 2.000, Polysorbate-20 2.800, microcryst. cellulose 116.800, NaCl 228.000, Povidone 60.000, PEG 160.000, HPMC-2208 14.000, colloidal SiO<sub>2</sub> 7.600, and Mg. The coating formulation also contained risperidone 5.000 mg.

IT 2058-52-8, Clothiapine 111974-69-7, Quetiapine

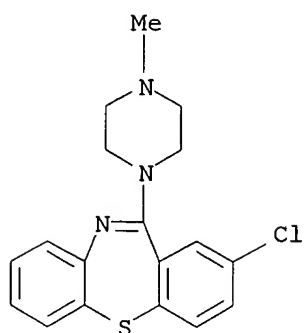
RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(osmotic device contg. alprazolam and antipsychotic agent)

RN 2058-52-8 CAPLUS

CN Dibenzo[b,f][1,4]thiazepine, 2-chloro-11-(4-methyl-1-piperazinyl)- (7CI, 8CI, 9CI) (CA INDEX NAME)

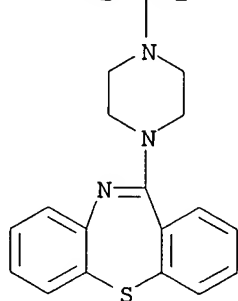
10/009,574



RN 111974-69-7 CAPLUS

CN Ethanol, 2-[2-(4-dibenzo[b,f][1,4]thiazepin-11-yl-1-piperazinyl)ethoxy]-  
(9CI) (CA INDEX NAME)

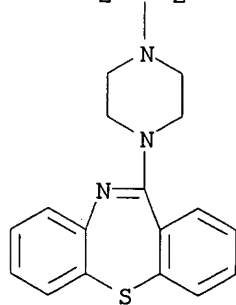
HO-CH<sub>2</sub>-CH<sub>2</sub>-O-CH<sub>2</sub>-CH<sub>2</sub>



RE.CNT 2 THERE ARE 2 CITED REFERENCES AVAILABLE FOR THIS RECORD  
ALL CITATIONS AVAILABLE IN THE RE FORMAT

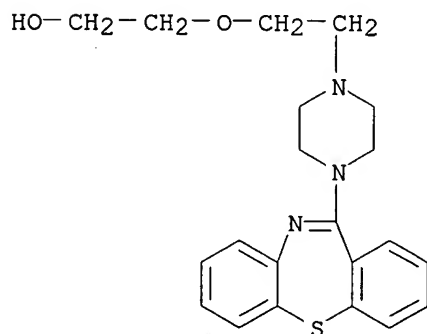
~~D30~~ ANSWER 18 OF 43 CAPLUS COPYRIGHT 2003 ACS  
~~AN~~ 2001:82838 CAPLUS  
~~DN~~ 135:131527  
 TI Antipsychotic treatment of **psychosis** and agitation in the elderly  
 AU Daniel, David G.  
 CS Department of Psychiatry and Behavioral Sciences, George Washington University, Washington, DC, USA  
 SO Journal of Clinical Psychiatry (2000), 61(Suppl. 14), 49-52  
 CODEN: JCLPDE; ISSN: 0160-6689  
 PB Physicians Postgraduate Press, Inc.  
 DT Journal; General Review  
 LA English  
 AB A review with 33 refs. Agitated, aggressive behavior and **psychosis** are common manifestations of Alzheimer's disease that frequently lead to institutionalization. The usefulness of conventional neuroleptic treatment in this population is limited by narrow therapeutic windows because of limited efficacy and high sensitivity to side effects. More recently, investigational clin. trials have suggested potential utility for atypical antipsychotics such as risperidone, olanzapine, and quetiapine in treatment of behaviorally disturbed individuals and for the psychotic manifestations of dementia.  
 IT 111974-69-7, Quetiapine  
 RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (antipsychotic treatment of **psychosis** and agitation in elderly humans)  
 RN 111974-69-7 CAPLUS  
 CN Ethanol, 2-[2-(4-dibenzo[b,f][1,4]thiazepin-11-yl-1-piperazinyl)ethoxy]-(9CI) (CA INDEX NAME)

HO-CH<sub>2</sub>-CH<sub>2</sub>-O-CH<sub>2</sub>-CH<sub>2</sub>



RE.CNT 33      THERE ARE 33 CITED REFERENCES AVAILABLE FOR THIS RECORD  
 ALL CITATIONS AVAILABLE IN THE RE FORMAT

~~130~~ ANSWER 19 OF 43 CAPLUS COPYRIGHT 2003 ACS  
~~AN~~ 2001:82837 CAPLUS  
~~DN~~ 135:131526  
 TI New treatments for bipolar disorder: The role of atypical neuroleptic agents  
 AU Ghaemi, S. Nassir  
 CS Consolidated Department of Psychiatry, Harvard Medical School, Psychopharmacology Program, Cambridge Hospital, Cambridge, MA, 02139, USA  
 SO Journal of Clinical Psychiatry (2000), 61(Suppl. 14), 33-42  
 CODEN: JCLPDE; ISSN: 0160-6689  
 PB Physicians Postgraduate Press, Inc.  
 DT Journal; General Review  
 LA English  
 AB A review with 78 refs. Atypical neuroleptic agents are an excellent, safer, and more effective alternative to the widespread practice of maintenance adjunctive treatment with traditional neuroleptic agents in patients with bipolar disorder. Currently, a no. of prospective studies are available with clozapine, risperidone, olanzapine, and quetiapine in the treatment of bipolar disorder. Most are short-term studies, although longer-term data are becoming available. Four double-blind studies of acute mania have been conducted with risperidone and olanzapine, leading to recent Food and Drug Administration approval for olanzapine in the indication of acute mania. Given the limited longer-term data, and the evidence for mostly adjunctive benefits with these agents, it seems unlikely that these agents will prove to be primary mood stabilizers in their own right. Nonetheless, they serve an important role as adjunctive treatments along with std. mood stabilizers in the rational polypharmacy of bipolar disorder. To date, differences in efficacy have not been established. However, differences in the side effect of wt. gain may be even more relevant in bipolar disorder than in schizophrenia due to the need to use std. mood stabilizers that often potentiate such wt. gain.  
 IT 111974-69-7, Quetiapine  
 RL: ADV (Adverse effect, including toxicity); BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (new atypical neuroleptic agents in treatments of bipolar disorder in humans)  
 RN 111974-69-7 CAPLUS  
 CN Ethanol, 2-[2-(4-dibenzo[b,f][1,4]thiazepin-11-yl-1-piperazinyl)ethoxy]-(9CI) (CA INDEX NAME)



RE.CNT 78 THERE ARE 78 CITED REFERENCES AVAILABLE FOR THIS RECORD

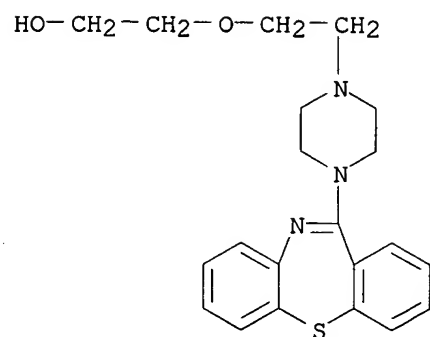


10/009,574

ALL CITATIONS AVAILABLE IN THE RE FORMAT

L30 ANSWER 20 OF 43 CAPLUS COPYRIGHT 2003 ACS  
AN 2001:70664 CAPLUS  
DN 135:132202  
TI The long-term effect of quetiapine (Seroquel) monotherapy on  
**weight** in patients with schizophrenia  
AU Brecher, M.; Rak, I. W.; Melvin, K.; Jones, A. M.  
CS AstraZeneca, Wilmington, DE, USA  
SO International Journal of Psychiatry in Clinical Practice (2000), 4(4),  
287-291  
CODEN: IJPCFZ; ISSN: 1365-1501  
PB Martin Dunitz Ltd.  
DT Journal  
LA English  
AB INTRODUCTION: Quetiapine (Seroquel) is an atypical antipsychotic drug with  
demonstrated efficacy and tolerability. In particular, placebo level  
extrapyramidal symptoms (EPS) across the entire dose range and a low  
propensity to cause sexual dysfunction suggest it may be assocd. with  
greater patient acceptability than alternative treatments. However, other  
side-effects, such as **wt.** gain, may also have a significant  
impact on treatment acceptability. METHOD: We report the long-term  
**wt.** changes obsd. in a cohort of 427 patients with schizophrenia  
from controlled and open-label extension (OLE) trials, in which quetiapine  
(mean dose 475 mg/day after 1 yr) was the only antipsychotic medication  
during the OLE period. RESULTS: In these patients, there was no overall  
effect on **wt.** across the body mass index (BMI) spectrum. There  
were no dose-related effects on **wt.**, and only one patient  
withdrew from treatment due to an adverse event of **wt.** gain.  
Quetiapine appeared to have a **wt.** neutral or 'normalizing'  
effect, with a tendency towards favorable shifts in bodyweight in  
underweight patients (BMI < 18.5 kg/m<sup>2</sup>) and severely obese patients (BMI  
.gtoreq. 35 kg/m<sup>2</sup>). CONCLUSION: These results indicate that long-term  
**wt.** changes with quetiapine monotherapy are minimal and  
potentially beneficial, and do not appear to raise the medical concerns  
assocd. with some other atypical agents.  
IT 111974-72-2, Seroquel  
RL: ADV (Adverse effect, including toxicity); BAC (Biological activity or  
effector, except adverse); BSU (Biological study, unclassified); THU  
(Therapeutic use); BIOL (Biological study); USES (Uses)  
(long-term effect of quetiapine (Seroquel) monotherapy on **wt.**  
in humans with schizophrenia)  
RN 111974-72-2 CAPLUS  
CN Ethanol, 2-[2-(4-dibenzo[b,f][1,4]thiazepin-11-yl-1-piperazinyl)ethoxy]-,  
(2E)-2-butenedioate (2:1) (salt) (9CI) (CA INDEX NAME)  
CM 1  
CRN 111974-69-7  
CMF C21 H25 N3 O2 S

10/009,574

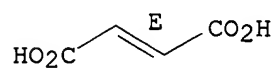


CM 2

CRN 110-17-8

CMF C4 H4 O4

Double bond geometry as shown.



RE.CNT 23 THERE ARE 23 CITED REFERENCES AVAILABLE FOR THIS RECORD  
ALL CITATIONS AVAILABLE IN THE RE FORMAT

~~130~~ ANSWER 21 OF 43 CAPLUS COPYRIGHT 2003 ACS  
~~AN~~ 2000:881023 CAPLUS  
~~DN~~ 134:33017

TI Combination for treating **weight** gain associated with  
antipsychotic use comprising an atypical antipsychotic and an H2  
antagonist

IN Todd, Jane Rogers

PA Eli Lilly and Company, USA

SO PCT Int. Appl., 43 pp.

CODEN: PIXXD2

DT Patent

LA English

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 2000074784	A1	20001214	WO 2000-US9811	20000522
	W:	AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM			
	RW:	GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG			
	EP 1189662	A1	20020327	EP 2000-931932	20000522
	R:	AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO			

PRAI US 1999-138315P P 19990609  
WO 2000-US9811 W 20000522

AB The invention provides methods and compns. for the prevention and treatment of **wt.** gain assocd. with antipsychotic use. These methods and compns. employ a compd. having activity as an atypical antipsychotic and an H2 antagonist. A capsule contained olanzapine 25, nizatidine 150, starch 150, and Mg stearate 210 mg.

IT **111974-69-7**, Quetiapine

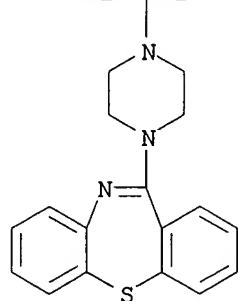
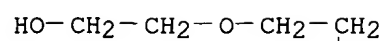
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(atypical antipsychotic and H2 antagonist combination for treating **wt.** gain assocd. with antipsychotic therapy)

RN 111974-69-7 CAPLUS

CN Ethanol, 2-[2-(4-dibenzo[b,f][1,4]thiazepin-11-yl-1-piperazinyl)ethoxy]-(9CI) (CA INDEX NAME)

10/009,574



RE.CNT 7      THERE ARE 7 CITED REFERENCES AVAILABLE FOR THIS RECORD  
ALL CITATIONS AVAILABLE IN THE RE FORMAT

~~LSO~~ ANSWER 22 OF 43 CAPLUS COPYRIGHT 2003 ACS

~~PN~~ 2000:841965 CAPLUS

~~DN~~ 134:535

TI Method of treatment

IN Reinstein, Michael J.; Jones, Andrew Martin

PA Astrazeneca AB, Swed.

SO PCT Int. Appl., 9 pp.

CODEN: PIXXD2

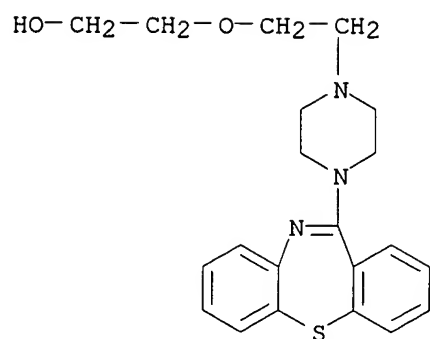
DT Patent

LA English

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 2000071106	A2	20001130	WO 2000-GB1875	20000516
	WO 2000071106	A3	20020510		
	W:				
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	CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU,				
	ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU,				
	LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE,				
	SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA,				
	ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
	RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE,				
	DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF,				
	CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				
EP	1223939	A1	20020724	EP 2000-927593	20000516
	R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,				
	IE, SI, LT, LV, FI, RO, MK, CY, AL				
JP	2003500353	T2	20030107	JP 2000-619413	20000516
PRAI	GB 1999-11499	A	19990519		
	GB 2000-2762	A	20000208		
	WO 2000-GB1875	W	20000516		
AB	A method of treating wt. in patients, in particular those suffering from <b>psychoses</b> , by administering the antipsychotic agent quetiapine.				
IT	<b>111974-72-2</b> , Quetiapine fumarate				
	RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)				
	(treatment of wt. gain in patients with antipsychotic quetiapine)				
RN	111974-72-2 CAPLUS				
CN	Ethanol, 2-[2-(4-dibenzo[b,f][1,4]thiazepin-11-yl-1-piperazinyl)ethoxy]-, (2E)-2-butenedioate (2:1) (salt) (9CI) (CA INDEX NAME)				
CM	1				
CRN	111974-69-7				
CMF	C21 H25 N3 O2 S				

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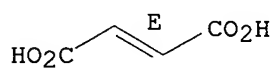


CM 2

CRN 110-17-8

CMF C4 H4 O4

Double bond geometry as shown.



L30 ANSWER 23 OF 43 CAPLUS COPYRIGHT 2003 ACS  
 AN 2000:773298 CAPLUS  
 DN 134:361239  
 TI Long-term use of quetiapine in elderly patients with psychotic disorders  
 AU Tariot, Pierre N.; Salzman, Carl; Yeung, Paul P.; Pultz, Joseph; Rak, Ihor W.  
 CS University of Rochester School of Medicine, Rochester, NY, USA  
 SO Clinical Therapeutics (2000), 22(9), 1068-1084  
 CODEN: CLTHDG; ISSN: 0149-2918  
 PB Excerpta Medica, Inc.  
 DT Journal  
 LA English  
 AB Quetiapine is an atypical antipsychotic agent that does not appear to increase patient risk for treatment-emergent extrapyramidal symptoms (EPS) or anticholinergic symptoms. Previous studies of quetiapine use in elderly patients with schizophrenia and other **psychoses** examd. short-term administration (.ltoreq.12 wk). Given the growing elderly population, the commensurate increase in elderly patients with **psychoses**, and the expected increase in disease treatment-years, the effect of long-term quetiapine administration in older patients is of considerable interest. This study assesses the long-term tolerability, safety, and clin. benefit of quetiapine in elderly patients with **psychosis**. Elderly patients (.gtoreq.65 yr of age) with psychotic disorders, as defined by the Diagnostic and Statistical Manual of Mental Disorders, Fourth Edition, participated in this 52-wk, open-label, multicenter trial. Investigators increased (and later adjusted) daily doses of quetiapine on the basis of clin. response and tolerability, and assessed safety and efficacy. Efficacy assessments were made using the 18-item Brief Psychiatric Rating Scale (BPRS), Clin. Global Impressions (CGI), Simpson-Angus Scale, and the Abnormal Involuntary Movement Scale (AIMS). For patients who withdrew before week 52, analyses were performed using obsd. data and the last observation carried forward. One hundred eighty-four patients with psychotic disorders (98 women and 86 men) with a mean age of 76.1 yr entered the trial. Seventy-two percent had psychotic disorders due to general medical conditions such as Alzheimer's disease, and 28% had other psychotic disorders, most commonly schizophrenia. Overall, 89 (48%) patients completed treatment through 52 wk. Median total daily dose was 137.5 mg. Reasons for withdrawal included lack of efficacy (19%), adverse events or intercurrent illness (15%), failure to return for follow-up (13%), protocol noncompliance (3%), and diminished need for treatment (2%). Somnolence (31%), dizziness (17%), and postural hypotension (15%) were common adverse events, but they rarely resulted in withdrawal from. Therapy. EPS-related adverse events occurred in 13% of patients. At end point (week 52), mean total score on the Simpson-Angus Scale had decreased from baseline by 1.8 points, whereas changes in AIMS scores were negligible. No clin. important effects were reported relative to mean changes in hematol., thyroid function, or hepatic function variables. Quetiapine treatment appeared to have no assocd. cardiovascular adverse outcomes despite cardiovascular comorbidities and unrestricted use of concomitant cardiovascular medications. Significant decreases in BPRS total score (n = 170, P < 0.001) and CGI Severity of Illness item score (n = 177, P < 0.002) were seen at end point (obsd. data and last observation carried forward). Decreases of .gtoreq.20% in mean BPRS total score were obsd. in 83 (49%) patients. These results provide preliminary information to clinicians regarding tolerability, safety, and clin. improvement with quetiapine in elderly patients with psychotic symptoms, and support controlled studies of quetiapine in this patient population.



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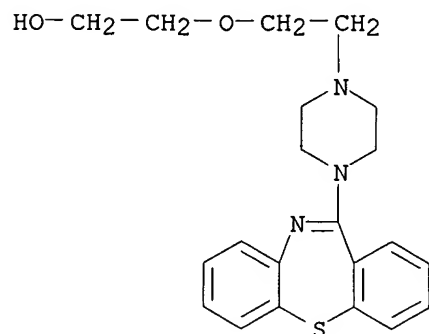
IT 111974-69-7, Quetiapine

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(long-term use of quetiapine in elderly patients with psychotic disorders)

RN 111974-69-7 CAPLUS

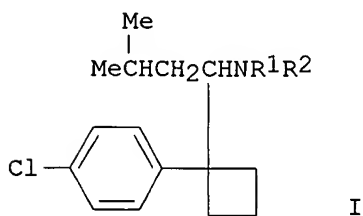
CN Ethanol, 2-[2-(4-dibenzo[b,f][1,4]thiazepin-11-yl-1-piperazinyl)ethoxy]-(9CI) (CA INDEX NAME)



RE.CNT 36 THERE ARE 36 CITED REFERENCES AVAILABLE FOR THIS RECORD  
ALL CITATIONS AVAILABLE IN THE RE FORMAT

L30 ANSWER 24 OF 43 CAPLUS COPYRIGHT 2003 ACS  
 AN 2000:688070 CAPLUS  
 DN 133:232860  
 TI Sibutramine and N-demethyl derivatives thereof for controlling  
**weight** gain associated with therapeutic drugs  
 IN Mendel, Carl M.; Seaton, Timothy B.; Weinstein, Steve P.  
 PA Knoll Pharmaceutical Company, USA  
 SO PCT Int. Appl., 17 pp.  
 CODEN: PIXXD2  
 DT Patent  
 LA English  
 FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 2000056313	A1	20000928	WO 2000-US7130	20000317
	W: AT, AU, BG, BR, CA, CN, CZ, DE, DK, ES, FI, GB, HR, HU, ID, IL, IN, IS, JP, KR, LT, LU, LV, MX, NO, NZ, PL, PT, RO, RU, SE, SG, SI, SK, TR, UA, ZA				
	RW: AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE				
	NZ 514009	A	20010928	NZ 2000-514009	20000317
	EP 1162965	A1	20011219	EP 2000-916480	20000317
	R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO				
	BR 2000009159	A	20011226	BR 2000-9159	20000317
	US 6376552	B1	20020423	US 2000-527962	20000317
	JP 2002539251	T2	20021119	JP 2000-606218	20000317
	NO 2001004480	A	20011102	NO 2001-4480	20010914
	BG 105997	A	20020628	BG 2001-105997	20011010
PRAI	US 1999-125340P	P	19990319		
	WO 2000-US7130	W	20000317		
OS	MARPAT 133:232860				
GI					

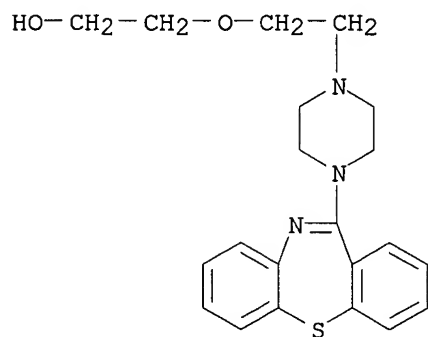


AB Compds. I (R1, R2 = H, Me) or a pharmaceutically acceptable salt thereof (e.g. N,N,-dimethyl-1-[1-(4-chlorophenyl)cyclobutyl]-3-methylbutylamine-HCl, optionally in the form of its monohydrate) are used for treating **wt.** gain assocd. with drug therapy, including the use of tricyclic antidepressants, lithium, sulfonylureas, .beta.-adrenergic blockers, certain steroid contraceptives, corticosteroids, insulin, cyproheptadine, sodium valproate, neuroleptics, phenothiazines, or piztifin.

IT **111974-69-7, Quetiapine**  
 RL: ADV (Adverse effect, including toxicity); BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (sibutramine and N-demethyl derivs. for controlling **wt.** gain

10/009,574

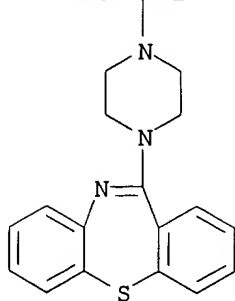
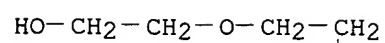
          assocd. with drug therapy)  
RN    111974-69-7    CAPLUS  
CN    Ethanol, 2-[2-(4-dibenzo[b,f][1,4]thiazepin-11-yl-1-piperazinyl)ethoxy]-  
      (9CI)    (CA INDEX NAME)



RE.CNT 2       THERE ARE 2 CITED REFERENCES AVAILABLE FOR THIS RECORD  
          ALL CITATIONS AVAILABLE IN THE RE FORMAT

L30 ANSWER 25 OF 43 CAPLUS COPYRIGHT 2003 ACS  
AN 2000:671544 CAPLUS  
DN 134:172545  
TI Quetiapine: A review of its clinical potential in the management of  
psychotic symptoms in Parkinson's disease  
AU Matheson, Anna J.; Lamb, Harriet M.  
CS Adis International Limited, Auckland, N. Z.  
SO CNS Drugs (2000), 14(2), 157-172  
CODEN: CNDREF; ISSN: 1172-7047  
PB Adis International Ltd.  
DT Journal; General Review  
LA English  
AB A review with 54 refs. Quetiapine is a dibenzothiazepine atypical  
antipsychotic which has a close pharmacol. resemblance to clozapine. In a  
no. of small noncomparative clin. trials, quetiapine has been successfully  
used in the treatment of **psychosis** in patients with Parkinson's  
disease. **Psychosis** in these patients is caused by current  
antiparkinsonian drug therapy, the underlying disease pathol. or a  
combination of both factors. In patients with Parkinson's disease with or  
without previous exposure to antipsychotics, quetiapine reduced psychotic  
symptoms as measured by a redn. in Brief Psychiatric Rating Scale scores  
from baseline. Quetiapine was also effective after treatment failure with  
clozapine, risperidone or olanzapine, and in psychiatrically stable  
patients who were switched from either clozapine or olanzapine. Motor  
function was generally maintained in most patients. In 2 of the largest  
trials, patients with Parkinson's disease reported adverse events such as  
headache, nausea, orthostatic hypotension, dizziness and diarrhoea after  
initiation of quetiapine therapy. In two 12-mo trials no development or  
exacerbation of extrapyramidal symptoms (EPS) occurred after the  
initiation of quetiapine therapy in patients with Parkinson's disease. In  
another trial, EPS were reported in 3% of patients with Parkinson's  
disease given quetiapine after treatment failure with another atypical  
antipsychotic. The incidence of EPS was generally not significantly  
different between quetiapine (75 to 750 mg/day) and placebo in patients  
with schizophrenia. If dosage redn. of antiparkinsonian therapy does not  
alleviate psychotic symptoms in patients with Parkinson's disease,  
quetiapine may offer an effective alternative to other atypical  
antipsychotic agents, without compromising motor function. Confirmation  
of the relative efficacy and low EPS potential of quetiapine in  
comparative trials with other atypical agents would be beneficial.  
However, based on the available data quetiapine is a treatment option for  
the management of this difficult-to-treat patient group.  
IT 111974-69-7, Quetiapine  
RL: ADV (Adverse effect, including toxicity); BAC (Biological activity or  
effector, except adverse); BSU (Biological study, unclassified); THU  
(Therapeutic use); BIOL (Biological study); USES (Uses)  
(quetiapine and its clin. potential in management of psychotic symptoms  
in Parkinson's disease)  
RN 111974-69-7 CAPLUS  
CN Ethanol, 2-[2-(4-dibenzo[b,f][1,4]thiazepin-11-yl-1-piperazinyl)ethoxy]-  
(9CI) (CA INDEX NAME)

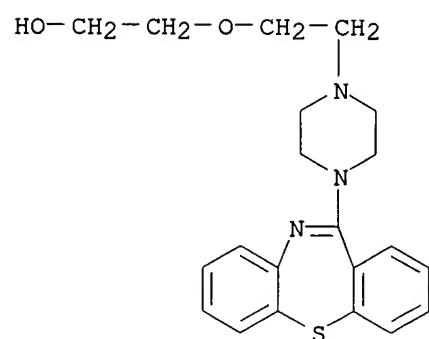
10/009,574



RE.CNT 54      THERE ARE 54 CITED REFERENCES AVAILABLE FOR THIS RECORD  
ALL CITATIONS AVAILABLE IN THE RE FORMAT

L30 ANSWER 26 OF 43 CAPLUS COPYRIGHT 2003 ACS  
AN 2000:432653 CAPLUS  
DN 133:37510  
TI Review of quetiapine and its clinical applications in schizophrenia  
AU Kasper, Siegfried; Muller-Spahn, Franz  
CS Department of General Psychiatry, University of Vienna, Vienna, Austria  
SO Expert Opinion on Pharmacotherapy (2000), 1(4), 783-801  
CODEN: EOPHF7; ISSN: 1465-6566  
PB Ashley Publications Ltd.  
DT Journal; General Review  
LA English  
AB A review with 87 refs. Preclin. studies have shown that quetiapine (Seroquel, AstraZeneca) is an atypical antipsychotic with many similarities to clozapine. Both placebo-controlled and comparative studies in patients with schizophrenia have demonstrated that quetiapine has long-term efficacy in both pos. and neg. domains, as well as beneficial effects on affective and cognitive symptoms. Comparative clin. studies confirm that quetiapine is at least as effective as the std. antipsychotics, chlorpromazine and haloperidol and response rates with quetiapine are similar to those reported with other atypical antipsychotics. Quetiapine has also demonstrated superior efficacy to haloperidol in partially responsive patients, who can be particularly difficult to treat. Quetiapine has a wide clin. dosing range (150-750 mg/day), although doses of 400 mg or above should be used in patients who do not fully respond to lower doses of the drug. Quetiapine is generally well tolerated with no requirement for routine ECG or blood monitoring and it has minimal effects on wt. Uniquely among other first-line atypical antipsychotics, quetiapine is assocd. with a placebo-level incidence of EPS and an indistinguishable effect from placebo on plasma prolactin at all doses. Thus, clinicians can confidently increase the dose of quetiapine, without increasing the risk of EPS or hyperprolactinemia. A no. of studies have also shown that quetiapine is well-tolerated and effective in patients who are particularly susceptible to EPS, including elderly and adolescent patients and those with pre-existing dopaminergic pathol., such as Alzheimer's disease and Parkinson's disease. The consistent efficacy in treating all schizophrenic domains and good tolerability, particularly placebo-level EPS, make quetiapine acceptable to patients, as demonstrated in a survey of patient satisfaction. Thus quetiapine is a suitable first-line therapy for the treatment of schizophrenia and **psychosis**.  
IT 111974-69-7, Quetiapine  
RL: ADV (Adverse effect, including toxicity); BAC (Biological activity or effector, except adverse); BPR (Biological process); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses)  
(quetiapine and its clin. applications in schizophrenia)  
RN 111974-69-7 CAPLUS  
CN Ethanol, 2-[2-(4-dibenzo[b,f][1,4]thiazepin-11-yl-1-piperazinyl)ethoxy]-(9CI) (CA INDEX NAME)

10/009,574

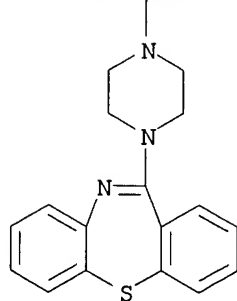
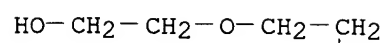


RE.CNT 88      THERE ARE 88 CITED REFERENCES AVAILABLE FOR THIS RECORD  
ALL CITATIONS AVAILABLE IN THE RE FORMAT

L30 ANSWER 27 OF 43 CAPLUS COPYRIGHT 2003 ACS  
AN 2000:297185 CAPLUS  
DN 132:303405  
TI Clinical predictors of acute response with quetiapine in psychotic mood disorders  
AU Zarate, Carlos A., Jr.; Rothschild, Anthony; Fletcher, Kenneth E.; Madrid, Alex; Zapatel, Jorge  
CS Bipolar and Psychotic Disorders Program and the Pharmacologic Research and Treatment Center, University of Massachusetts Medical School, Worcester, MA, 01655, USA  
SO Journal of Clinical Psychiatry (2000), 61(3), 185-189  
CODEN: JCLPDE; ISSN: 0160-6689  
PB Physicians Postgraduate Press, Inc.  
DT Journal  
LA English  
AB Background: In controlled studies of patients with schizophrenia, the atypical antipsychotic quetiapine, 300 mg/day, has been shown to be as effective in the treatment of pos. and neg. symptoms as haloperidol. However, little is known about the efficacy of quetiapine in patients with psychotic mood disorders. The purpose of this study was to assess the efficacy of quetiapine in the treatment of psychotic mood disorders in comparison with nonaffective psychotic disorders and identify clin. factors assocd. with quetiapine response. Method: In a naturalistic setting, by reviewing medical records, we assessed response to quetiapine and factors assocd. with response to quetiapine in 145 consecutive patients newly treated with the drug at a nonprofit academic psychiatric hospital. These patients had received a discharge diagnosis of bipolar disorder (manic, mixed, or depressive type), major depression with psychotic features, schizophrenia, schizoaffective disorder (bipolar or depressive type), delusional disorder, or **psychosis** not otherwise specified (NOS) according to DSM-IV criteria. Results: Patients with a diagnosis of bipolar disorder, manic, mixed, or depressed and schizoaffective disorder, bipolar type displayed higher response rates (> 74%) compared with patients with schizophrenia. However, this finding did not achieve statistical significance. A diagnosis of major depression with psychotic features ( $p = .02$ ) and longer duration of illness ( $p = .03$ ) were assocd. with less chance of responding. Conclusion: Quetiapine may be a useful alternative or adjunctive treatment for patients with bipolar and schizoaffective disorders.  
IT **111974-69-7, Quetiapine**  
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(clin. predictors of acute response to quetiapine in human psychotic mood disorders)  
RN 111974-69-7 CAPLUS  
CN Ethanol, 2-[2-(4-dibenzo[b,f][1,4]thiazepin-11-yl-1-piperazinyl)ethoxy]-(9CI) (CA INDEX NAME)



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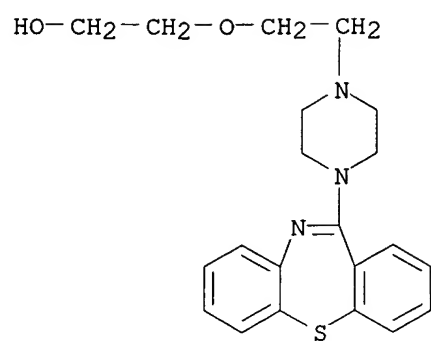


RE.CNT 18      THERE ARE 18 CITED REFERENCES AVAILABLE FOR THIS RECORD  
ALL CITATIONS AVAILABLE IN THE RE FORMAT

L30 ANSWER 28 OF 43 CAPLUS COPYRIGHT 2003 ACS  
 AN 2000:277835 CAPLUS  
 DN 132:298845  
 TI Therapy for improving cognition  
 IN De Nijs, Paul Leonce Irma; Parys, Wim Louis Julien  
 PA Janssen Pharmaceutica N.V., Belg.  
 SO PCT Int. Appl., 7 pp.  
 CODEN: PIXXD2  
 DT Patent  
 LA English  
 FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 2000023057	A2	20000427	WO 1999-EP7804	19991012
	WO 2000023057	A3	20000727		
	W:	AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM			
	RW:	GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG			
	CA 2345767	AA	20000427	CA 1999-2345767	19991012
	AU 9964727	A1	20000508	AU 1999-64727	19991012
	BR 9914419	A	20010626	BR 1999-14419	19991012
	EP 1121131	A2	20010808	EP 1999-952580	19991012
	R:	AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO			
	EE 200100136	A	20020617	EE 2001-136	19991012
	JP 2002527469	T2	20020827	JP 2000-576832	19991012
	BG 105302	A	20011130	BG 2001-105302	20010301
	NO 2001001403	A	20010320	NO 2001-1403	20010320
PRAI	EP 1998-203454	A	19981016		
	WO 1999-EP7804	W	19991012		
AB	The present invention is concerned with pharmaceutical compns. comprising a carrier and as first active ingredient an atypical antipsychotic agent (I) and as second active ingredient an acetylcholinesterase inhibitor (II), each in an amt. producing a therapeutically beneficial effect in patients suffering from <b>psychosis</b> , or Alzheimer's disease or related dementias. The therapeutically beneficial effect can be a synergistic effect on the cognitive functioning of patients suffering from Alzheimer's disease or related dementias or the prevention of the further deterioration of cognition in the patients, or the redn. of adverse effects assocd. with one of the active ingredients by the other of the active ingredients. Preferred compns. comprise risperidone as the atypical antipsychotic and galantamine as the acetylcholinesterase inhibitor.				
IT	111974-69-7, Quetiapine				
	RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)				
	(therapeutics for improving cognition contg. antipsychotic agent and acetylcholinesterase inhibitor)				
RN	111974-69-7 CAPLUS				
CN	Ethanol, 2-[2-(4-dibenzo[b,f][1,4]thiazepin-11-yl-1-piperazinyl)ethoxy]-(9CI) (CA INDEX NAME)				

10/009,574



L30 ANSWER 29 OF 43 CAPLUS COPYRIGHT 2003 ACS

AN 2000:260000 CAPLUS

DN 132:288772

TI Use of metformin to counteract **weight** gain associated with  
valproate and other psychotropic medications

IN Cottingham, Elizabeth Marie

PA Children's Hospital Research Foundation, USA; Morrison, John Ainslie

SO PCT Int. Appl., 14 pp.

CODEN: PIXXD2

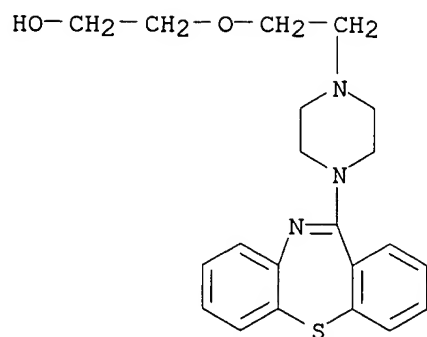
DT Patent

LA English

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 2000021522	A1	20000420	WO 1999-US24262	19991015
	W:	AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM			
	RW:	GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG			
	US 6194466	B1	20010227	US 1999-416330	19991012
	AU 9964328	A1	20000501	AU 1999-64328	19991015
	EP 1121110	A1	20010808	EP 1999-952021	19991015
	R:	AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO			
PRAI	US 1998-104394P	P	19981015		
	US 1999-416330	A	19991012		
	WO 1999-US24262	W	19991015		
AB	A method for minimizing the <b>wt.</b> gain side effect assocd. with psychotropic treatment is disclosed. In the method, Metformin, a biguanide compd., is concurrently administered to a patient taking the psychotropic active. A pharmaceutical compn. contg. the combination of psychotropic active and Metformin is also disclosed. Psychotropic actives are selected from valproate, Risperdal, Lithobid, Zyprexa and Seroquel.				
IT	<b>111974-72-2</b> , Seroquel				
	RL: ADV (Adverse effect, including toxicity); BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)				
	(metformin to counteract <b>wt.</b> gain assocd. with valproate and other psychotropic medications)				
RN	111974-72-2 CAPLUS				
CN	Ethanol, 2-[2-(4-dibenzo[b,f][1,4]thiazepin-11-yl-1-piperazinyl)ethoxy]-, (2E)-2-butenedioate (2:1) (salt) (9CI) (CA INDEX NAME)				
CM	1				
CRN	111974-69-7				
CMF	C21 H25 N3 O2 S				

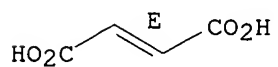
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CM. 2

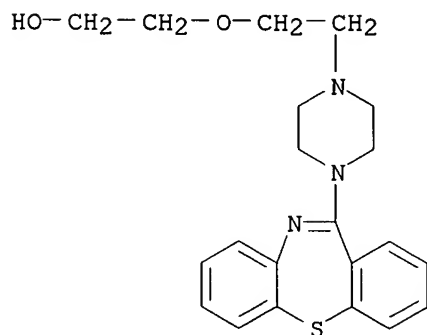
CRN 110-17-8  
CMF C4 H4 O4

Double bond geometry as shown.



RE.CNT 8      THERE ARE 8 CITED REFERENCES AVAILABLE FOR THIS RECORD  
ALL CITATIONS AVAILABLE IN THE RE FORMAT

L30 ANSWER 30 OF 43 CAPLUS COPYRIGHT 2003 ACS  
 AN 2000:243330 CAPLUS  
 DN 132:260034  
 TI The efficacy of atypical antipsychotics in the treatment of depressive symptoms, hostility, and suicidality in patients with schizophrenia  
 AU Keck, Paul E., Jr.; Strakowski, Stephen M.; McElroy, Susan L.  
 CS Biological Psychiatry and Psychotic Disorders Research Programs, Department of Psychiatry, University of Cincinnati College of Medicine, Cincinnati, OH, 45267-0559, USA  
 SO Journal of Clinical Psychiatry (2000), 61(Suppl. 3), 4-9  
 CODEN: JCLPDE; ISSN: 0160-6689  
 PB Physicians Postgraduate Press, Inc.  
 DT Journal; General Review  
 LA English  
 AB A review with 80 refs. Depressive symptoms and syndromal depression commonly occur in patients with schizophrenia. Schizophrenia is also assocd. with aggression directed at self and others. For this article, the available literature regarding the efficacy of clozapine, risperidone, olanzapine, quetiapine, and ziprasidone in the treatment of depression, hostility, and suicidality in patients with schizophrenia was reviewed. These studies suggest that atypical antipsychotics may exert therapeutic effects on depression and hostility as well as **psychosis** and that clozapine and olanzapine may reduce suicidality in patients with schizophrenia. These therapeutic actions appear to represent addnl. advantages of atypical antipsychotics compared with std. agents.  
 IT **111974-69-7, Quetiapine**  
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (efficacy of atypical antipsychotics in treatment of depressive symptoms, hostility, and suicidality in patients with schizophrenia)  
 RN 111974-69-7 CAPLUS  
 CN Ethanol, 2-[2-(4-dibenzo[b,f][1,4]thiazepin-11-yl-1-piperazinyl)ethoxy]-(9CI) (CA INDEX NAME)



RE.CNT 80 THERE ARE 80 CITED REFERENCES AVAILABLE FOR THIS RECORD  
 ALL CITATIONS AVAILABLE IN THE RE FORMAT

~~DSO~~ ANSWER 31 OF 43 CAPLUS COPYRIGHT 2003 ACS

~~AN~~ 2000:241564 CAPLUS

~~DN~~ 132:288780

TI Methods of identifying inverse agonists of the serotonin 2a receptor, therapeutic and diagnostic methods, and test kit

IN Weiner, David; Brann, Mark R.

PA Acadia Pharmaceuticals Inc., USA

SO PCT Int. Appl., 42 pp.

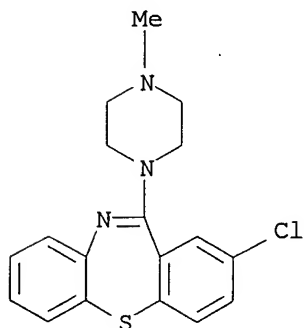
CODEN: PIXXD2

DT Patent

LA English

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 2000020636	A1	20000413	WO 1999-US21439	19991007
	W:	AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM			
	RW:	GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG			
	US 6358698	B1	20020319	US 1999-413626	19991006
	AU 9963912	A1	20000426	AU 1999-63912	19991007
PRAI	US 1998-103317P	P	19981007		
	US 1999-413626	A1	19991006		
	WO 1999-US21439	W	19991007		
AB	A method for identifying compds. which act as inverse agonists of the 5-HT <sub>2A</sub> receptor comprises contacting a constitutively active 5-HT <sub>2A</sub> receptor with at least one test compd. and detg. any decrease in the level of basal activity of the receptor. The inverse agonists may be used in the treatment of schizophrenia and related <b>psychoses</b> .				
IT	<b>2058-52-8</b> , Clothiapine <b>264256-90-8</b> , Quetiapine RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (serotonin 2a receptor inverse agonist identification, therapeutic and diagnostic methods, and test kit)				
RN	2058-52-8 CAPLUS				
CN	Dibenzo[b,f][1,4]thiazepine, 2-chloro-11-(4-methyl-1-piperazinyl)- (7CI, 8CI, 9CI) (CA INDEX NAME)				



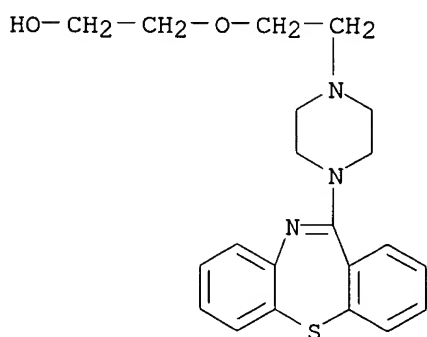
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RN 264256-90-8 CAPLUS

RE.CNT 7 THERE ARE 7 CITED REFERENCES AVAILABLE FOR THIS RECORD  
ALL CITATIONS AVAILABLE IN THE RE FORMAT

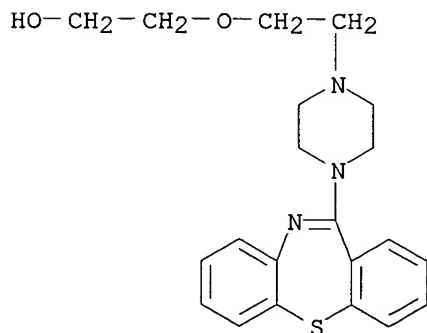


L30 ANSWER 32 OF 43 CAPLUS COPYRIGHT 2003 ACS  
 AN 2000:231510 CAPLUS  
 DN 132:231892  
 TI Switching outpatients between atypical antipsychotics  
 AU Bogan, Ann M.; Shellhorn, Eric; Brown, E. Sherwood; Mcdanald, Conway;  
 Suppes, Trisha  
 CS University of Texas Health Science Center, Houston, TX, USA  
 SO Progress in Neuro-Psychopharmacology & Biological Psychiatry (2000),  
 24(2), 351-355  
 CODEN: PNPPD7; ISSN: 0278-5846  
 PB Elsevier Science Inc.  
 DT Journal  
 LA English  
 AB Some reports have suggested an increase in symptoms when switching  
 patients with **psychosis** from clozapine to other atypical  
 antipsychotics. No data are available on switching between atypical  
 antipsychotics other than clozapine, though this is common in clin.  
 practice. Six patients with schizophrenia or schizo-affective disorder,  
 bipolar type were switched to quetiapine after finishing a clin. trial of  
 sertindole. During the observation period of two to ten weeks no subjects  
 worsened and one improved. Side effects were mild. These preliminary  
 data suggest that switching between some atypical agents may be well  
 tolerated. Larger controlled trials are needed to confirm this  
 observation.  
 IT **111974-69-7, Quetiapine**  
 RL: ADV (Adverse effect, including toxicity); BAC (Biological activity or  
 effector, except adverse); BSU (Biological study, unclassified); THU  
 (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (switching outpatients between atypical antipsychotics)  
 RN 111974-69-7 CAPLUS  
 CN Ethanol, 2-[2-(4-dibenzo[b,f][1,4]thiazepin-11-yl-1-piperazinyl)ethoxy]-  
 (9CI) (CA INDEX NAME)



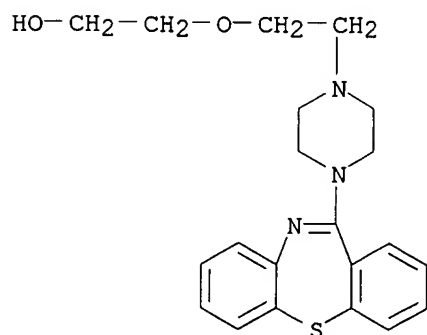
RE.CNT 6 THERE ARE 6 CITED REFERENCES AVAILABLE FOR THIS RECORD  
 ALL CITATIONS AVAILABLE IN THE RE FORMAT

L30 ANSWER 33 OF 43 CAPLUS COPYRIGHT 2003 ACS  
 AN 2000:173864 CAPLUS  
 DN 132:330032  
 TI New dopamine receptor, D2Longer, with unique TG splice site, in human brain  
 AU Seeman, P.; Nam, D.; Ulpian, C.; Liu, I. S. C.; Tallerico, T.  
 CS Department of Pharmacology, University of Toronto, Toronto, ON, Can.  
 SO Molecular Brain Research (2000), 76(1), 132-141  
 CODEN: MBREE4; ISSN: 0169-328X  
 PB Elsevier Science B.V.  
 DT Journal  
 LA English  
 AB Brain dopamine receptor agonists alleviate the signs of Parkinson's disease, while dopamine receptor antagonists alleviate hallucinations and delusions in **psychosis**. The dopamine type 2 receptor (or D2) is blocked by antipsychotic drugs, including even the "atypical" drugs such as clozapine or remoxipride, in direct relation to their clin. potencies. Compared to the long form of the D2 receptor (D2Long), the short form (D2Short) may be three times more sensitive to benzamide antipsychotic drugs. Hence, it is essential to identify addnl. variants of dopamine receptors for which more selective antipsychotic drugs can be found. Although no family linkage has been found between the D2 receptor and schizophrenia, there can be brain region abnormalities in the RNA transcript expression of dopamine receptors. Therefore, to identify variant dopamine D2 receptors, the authors searched for mutations in the RNA transcripts for the dopamine D2 receptor in the striatum of post-mortem brains from individuals who died with **psychosis**, including schizophrenia. A new splice variant of the D2 receptor, D2Longer, with a unique TG splice site, was found in one control brain and in two psychotic brains.  
 IT **111974-69-7**, Quetiapine  
 RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process)  
 (sequence and function of dopamine receptor D2Longer isoform from brain of psychotic and normal humans)  
 RN 111974-69-7 CAPLUS  
 CN Ethanol, 2-[2-(4-dibenzo[b,f][1,4]thiazepin-11-yl-1-piperazinyl)ethoxy]-(9CI) (CA INDEX NAME)



RE.CNT 50 THERE ARE 50 CITED REFERENCES AVAILABLE FOR THIS RECORD  
 ALL CITATIONS AVAILABLE IN THE RE FORMAT

L30 ANSWER 34 OF 43 CAPLUS COPYRIGHT 2003 ACS  
 AN 2000:97035 CAPLUS  
 DN 132:132262  
 TI Efficacy of quetiapine in Parkinson's patients with **psychosis**  
 AU Targum, Steven D.; Abbott, Jacob L.  
 CS Clinical Studies Limited, Philadelphia, PA, 19106, USA  
 SO Journal of Clinical Psychopharmacology (2000), 20(1), 54-60  
 CODEN: JCPYDR; ISSN: 0271-0749  
 PB Lippincott Williams & Wilkins  
 DT Journal  
 LA English  
 AB Eleven patients with Parkinson's disease (PD) and acute **psychosis** received flexible doses of quetiapine between 25 and 300 mg/day based on clin. response and tolerance. Ten patients were receiving dopaminergic agents at baseline. Serial efficacy ratings (Brief Psychiatric Rating Scale, Clin. Global Impressions Scale), neuromuscular symptom assessments (Abnormal Involuntary Movement Scale, Simpson-Angus Scale, Unified Parkinson's Disease Rating Scale [UPDRS]), and adverse events monitoring were performed for up to 52 wk. The patients had moderate hallucinations and/or delusions at baseline before the initiation of quetiapine. Nine of the 11 patients completed at least 12 wk of treatment. Quetiapine was well tolerated in all but one patient, who became dizzy within the first week and withdrew from the study. Ten patients presented with moderate visual hallucinations. Quetiapine was markedly effective in controlling visual hallucinations in six of these patients. Symptoms of paranoia or delusions were less responsive to quetiapine. Four patients withdrew because of adverse events or comorbid medical problems, two withdrew because of a lack of efficacy, and five completed 52 wk of treatment. The introduction of quetiapine did not exacerbate parkinsonian symptoms. Motor dysfunction, as measured by the UPDRS, revealed a slow, gradual worsening consistent with the progression of PD. Atypical antipsychotic medications such as quetiapine have a reduced likelihood of causing adverse drug-induced parkinsonism and therefore a possible role in treating psychotic symptoms in patients with PD.  
 IT **111974-69-7, Quetiapine**  
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (efficacy of quetiapine in Parkinson's patients with **psychosis**)  
 RN 111974-69-7 CAPLUS  
 CN Ethanol, 2-[2-(4-dibenzo[b,f][1,4]thiazepin-11-yl-1-piperazinyl)ethoxy]-(9CI) (CA INDEX NAME)



10/009,574

RE.CNT 25      THERE ARE 25 CITED REFERENCES AVAILABLE FOR THIS RECORD  
ALL CITATIONS AVAILABLE IN THE RE FORMAT

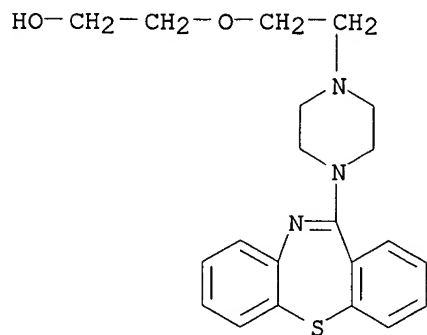
L30 ANSWER 35 OF 43 CAPLUS COPYRIGHT 2003 ACS  
AN 1999:582406 CAPLUS  
DN 131:208988  
TI Effect of clozapine-quetiapine combination therapy on **weight** and  
glycemic control: Preliminary findings  
AU Reinstein, Michael J.; Sirotovskaya, Larissa A.; Jones, Lynne E.; Mohan,  
Sangarapillai; Chasanov, Maxim A.  
CS Clinical Research Department, Forest Foundation Inc., Chicago, IL, USA  
SO Clinical Drug Investigation (1999), 18(2), 99-104  
CODEN: CDINFR; ISSN: 1173-2563  
PB Adis International Ltd.  
DT Journal  
LA English  
AB Objective: The purpose of this open-label, non-randomized, 10-mo,  
retrospective comparative study was to assess changes in **wt.** and  
**diabetes** status for patients initially treated with clozapine who  
developed **diabetes** and who were then switched to  
clozapine-quetiapine combination therapy. Methods: Sixty-five clinic  
charts were reviewed. All patients were from long-term care facilities.  
Bodyweight data were collected for this group of 65 randomly selected  
schizophrenic patients who were on clozapine initially (200 to 800 mg/day  
for 6 mo) and then had quetiapine (Seroquel) added to their therapy.  
Clozapine dosages were reduced as quetiapine was added proportionally: 25%  
of the clozapine dose was changed to quetiapine, using a ratio of exactly  
1mg clozapine to 2mg of quetiapine. The quetiapine dosages ranged from  
200 to 800 mg/day. This means that each patient received 6 mo of  
clozapine therapy followed by 10 mo of combination treatment with  
clozapine-quetiapine. **Wts.** were recorded monthly, and  
**diabetes** status was also performed for patients who developed the  
condition during clozapine monotherapy. Results: Changes in **wt.**  
and the status of **diabetes** were detd. in patients switched from  
a 6-mo clozapine therapy to the 10-mo combination clozapine-quetiapine  
treatment. All changes were statistically significant ( $p < 0.001$ ). Use  
of this combination therapy in the management of **wt.** gain and  
**diabetes** resulted in a 100% satisfactory response. All 65  
patients showed **wt.** loss ranging from 0.22 to 10.5kg (0.5 to  
23lb) [mean 1.8kg (3.98lb)] after the first month of combination therapy,  
and the improvement continued through the study duration (10 mo). Marked  
total **wt.** loss ranged from 0.45 to 18.6kg (1 to 41lb), with a  
mean loss of 4.2kg (9.2lb) over the 10-mo study period. 20% of patients  
(13 patients) who developed **diabetes** during the 6-mo clozapine  
monotherapy showed significant improvement of disease status with addn. of  
quetiapine. Compliance with medication was 100% and no significant  
adverse events were obsd. The most common adverse event reported by  
patients was drowsiness. However, this did not contribute a valid reason  
for discontinuation of clozapine-quetiapine therapy and could be cor. by  
dosage adjustment at any time of the report of this adverse effect by  
patients. Conclusion: An unexpected, yet welcome, clin. effect of  
quetiapine is its apparent propensity to induce **wt.** loss and  
improve glycemic control in patients who gain **wt.** and develop  
**diabetes** on clozapine therapy. The results of this retrospective  
study support the safety and tolerability of clozapine-quetiapine  
combination therapy.  
IT 111974-69-7, Quetiapine  
RL: ADV (Adverse effect, including toxicity); BAC (Biological activity or  
effector, except adverse); BSU (Biological study, unclassified); THU  
(Therapeutic use); BIOL (Biological study); USES (Uses)  
(effect of clozapine-quetiapine combination therapy on **wt.**

10/009,574

and glycemic control in schizophrenic humans)

RN 111974-69-7 CAPLUS

CN Ethanol, 2-[2-(4-dibenzo[b,f][1,4]thiazepin-11-yl-1-piperazinyl)ethoxy]-  
(9CI) (CA INDEX NAME)



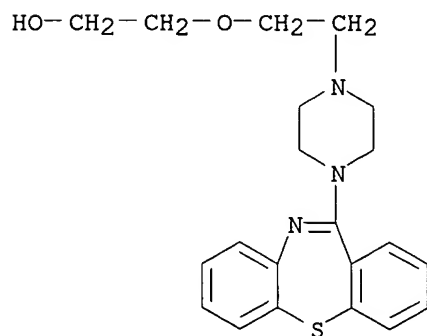
RE.CNT 10 THERE ARE 10 CITED REFERENCES AVAILABLE FOR THIS RECORD  
ALL CITATIONS AVAILABLE IN THE RE FORMAT

L30 ANSWER 36 OF 43 CAPLUS COPYRIGHT 2003 ACS  
 AN 1999:389659 CAPLUS  
 DN 131:39089  
 TI The role of atypical antipsychotics in the treatment of movement disorders  
 AU Fernandez, Hubert H.; Friedman, Joseph H.  
 CS Movement Disorders Unit, Department of Neurology, Memorial Hospital of  
 Rhode Island, Brown University School of Medicine, Pawtucket, RI, USA  
 SO CNS Drugs (1999), 11(6), 467-483  
 CODEN: CNDREF; ISSN: 1172-7047  
 PB Adis International Ltd.  
 DT Journal; General Review  
 LA English  
 AB A review with 190 refs. An atypical antipsychotic drug is loosely defined by its ability to produce an antipsychotic effect without inducing extrapyramidal symptoms (EPS). To date, 4 atypical antipsychotics have been released in the US: clozapine, quetiapine, olanzapine and risperidone, which are listed in decreasing order of "atypicality" based on clin. and preclin. studies. While the outcome of trials with quetiapine on parkinsonian patients (considered the most stringent test of the atypicality of a drug) is awaited, clozapine remains the prototypic atypical antipsychotic drug. Disappointing reports of risperidone-induced parkinsonism raise questions about the atypical nature of this drug. Olanzapine appears to be intermediate between risperidone and clozapine in inducing EPS. Drug-induced **psychosis** in Parkinson's disease and antipsychotic-induced movement disorders in psychotic patients are the most common indications for an atypical antipsychotic in patients with movement disorders. In drug-induced **psychosis** in Parkinson's disease, the antiparkinsonians are first reduced until the **psychosis** resolves. Unfortunately, motor function is often compromised as a result. The addn. of an atypical antipsychotic drug, without altering the regimen of antiparkinsonians, often controls **psychosis** without compromising motor function. Depending on the atypical antipsychotic used, the dosage required may be substantially lower than that for schizophrenic patients. No treatment strategy has been proven to be clearly superior in suppressing antipsychotic-induced movement disorders such as tardive dyskinesia, tardive akathisia and dystonia. Nonetheless, a review of the available data strongly suggests that clozapine has substantially less risk of inducing tardive dyskinesia than conventional antipsychotic agents. No case of tardive dyskinesia developing in patients who have taken clozapine as their only antipsychotic has yet been reported. Although there is evidence that clozapine may have an active therapeutic effect against pre-existing tardive dyskinesia, this remains inconclusive. Data on the use of clozapine for tremor in Parkinson's disease suggest significant benefit. Clozapine has also been reported to be useful in a variety of movement disorders including levodopa-induced dyskinesia, nocturnal akathisia and dystonia in Parkinson's disease, but the no. of patients involved is small. No definitive conclusion on the role of atypical antipsychotic agents in other behavioral disorders such as depression, anxiety and sleep fragmentation in Parkinson's disease, as well as in other movement disorders, can be made until well-planned long-term double-blind trials have been performed.  
 IT 111974-69-7, Quetiapine  
 RL: ADV (Adverse effect, including toxicity); BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (role of atypical antipsychotics in the treatment of human movement disorders)

10/009,574

RN 111974-69-7 CAPLUS

CN Ethanol, 2-[2-(4-dibenzo[b,f][1,4]thiazepin-11-yl-1-piperazinyl)ethoxy]-  
(9CI) (CA INDEX NAME)



RE.CNT 190 THERE ARE 190 CITED REFERENCES AVAILABLE FOR THIS RECORD  
ALL CITATIONS AVAILABLE IN THE RE FORMAT



~~130~~ ANSWER 37 OF 43 CAPLUS COPYRIGHT 2003 ACS

~~AN~~ 1998:527193 CAPLUS

~~DN~~ 129:166193

TI Therapeutic treatment and prevention of infections with a bioactive material encapsulated within a biodegradable-biocompatible polymeric matrix

IN Setterstrom, Jean A.; Van Hamont, John E.; Reid, Robert H.; Jacob, Elliot; Jeyanthi, Ramasubbu; Boedeker, Edgar C.; McQueen, Charles E.; Tice, Thomas R.; Roberts, F. Donald; Friden, Phil

PA United States Dept. of the Army, USA; Van Hamont, John E.; et al.

SO PCT Int. Appl., 363 pp.

CODEN: PIXXD2

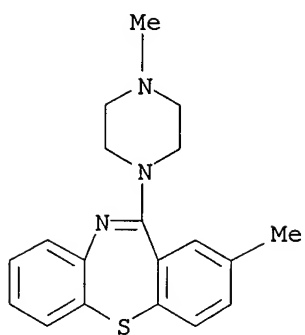
DT Patent

LA English

FAN.CNT 13

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 9832427	A1	19980730	WO 1998-US1556	19980127
	W:				
	AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GE, GH, HU, IL, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
	RW:				
	GH, GM, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG				
	US 6309669	B1	20011030	US 1997-789734	19970127
	AU 9863175	A1	19980818	AU 1998-63175	19980127
PRAI	US 1997-789734	A	19970127		
	US 1984-590308	B1	19840316		
	US 1992-867301	A2	19920410		
	US 1995-446148	A2	19950522		
	US 1995-446149	B2	19950522		
	US 1996-590973	B2	19960124		
	WO 1998-US1556	W	19980127		
AB	Novel burst-free, sustained release biocompatible and biodegradable microcapsules are disclosed which can be programmed to release their active core for variable durations ranging from 1-100 days in an aq. physiol. environment. The microcapsules are comprised of a core of polypeptide or other biol. active agent encapsulated in a matrix of poly(lactide/glycolide) copolymer, which may contain a pharmaceutically acceptable adjuvant, as a blend of uncapped free carboxyl end group and end-capped forms ranging in ratios from 100/0 to 1/99.				
IT	5800-19-1, Metiapine				
	RL: BPR (Biological process); BSU (Biological study, unclassified); DEV (Device component use); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses)				
	(prevention of infections with bioactive material encapsulated within biodegradable-biocompatible polymeric matrix)				
RN	5800-19-1 CAPLUS				
CN	Dibenzo[b,f][1,4]thiazepine, 2-methyl-11-(4-methyl-1-piperazinyl)- (7CI, 8CI, 9CI) (CA INDEX NAME)				

10/009,574



RE.CNT 6      THERE ARE 6 CITED REFERENCES AVAILABLE FOR THIS RECORD  
ALL CITATIONS AVAILABLE IN THE RE FORMAT

L30 ANSWER 38 OF 43 CAPLUS COPYRIGHT 2003 ACS

AN 1998:204419 CAPLUS

DN 128:261968

TI Pharmaceutical composition containing combination of atypical antipsychotic and serotonin reuptake inhibitor for treatment of **psychoses**

IN Bymaster, Franklin Porter; Perry, Kenneth Wayne; Tollefson, Gary Dennis

PA Eli Lilly and Co., USA

SO Eur. Pat. Appl., 15 pp.

CODEN: EPXXDW

DT Patent

LA English

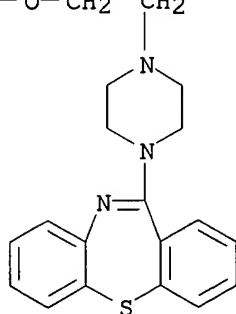
FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	EP 830864	A1	19980325	EP 1997-307375	19970922
	EP 830864	B1	20030129		
	R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO				
	ZA 9707967	A	19990304	ZA 1997-7967	19970904
	WO 9811897	A1	19980326	WO 1997-US15874	19970909
	W: AL, AM, AU, AZ, BA, BB, BG, BR, BY, CA, CN, CU, CZ, EE, GE, GH, HU, ID, IL, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, RO, RU, SD, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
	RW: GH, KE, LS, MW, SD, SZ, UG, ZW, BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG				
	AU 9744112	A1	19980414	AU 1997-44112	19970909
	AU 719033	B2	20000504		
	BR 9711530	A	19990824	BR 1997-11530	19970909
	CN 1230886	A	19991006	CN 1997-198113	19970909
	NZ 334168	A	20000929	NZ 1997-334168	19970909
	JP 2001503031	T2	20010306	JP 1998-514717	19970909
	EP 1256345	A1	20021113	EP 2002-16238	19970922
	R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, PT, IE, SI, LT, LV, FI, RO, AL				
	AT 231724	E	20030215	AT 1997-307375	19970922
	US 6147072	A	20001114	US 1997-935872	19970923
	NO 9901381	A	19990322	NO 1999-1381	19990322
	KR 2000048518	A	20000725	KR 1999-702422	19990322
PRAI	US 1996-26884P	P	19960923		
	WO 1997-US15874	W	19970909		
	EP 1997-307375	A3	19970922		
AB	Pharmaceutical compns. contg. combination of atypical antipsychotics and serotonin reuptake inhibitors are useful for the treatment of <b>psychoses</b> . Form II olanzapine (I) polymorph was prepd. by heating I at 76.degree. for 30 min in Et acetate and crystn. Hard gelatin capsules contained I 25, fluoxetine hydrochloride 20, starch 150, and magnesium stearate 10 mg.				
IT	<b>111974-69-7, Quetiapine</b>				
	RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)				
	(pharmaceutical compn. contg. combination of atypical antipsychotic and serotonin reuptake inhibitor for treatment of <b>psychoses</b> )				
RN	111974-69-7 CAPLUS				
CN	Ethanol, 2-[2-(4-dibenzo[b,f][1,4]thiazepin-11-yl-1-piperazinyl)ethoxy]-				

10/009,574

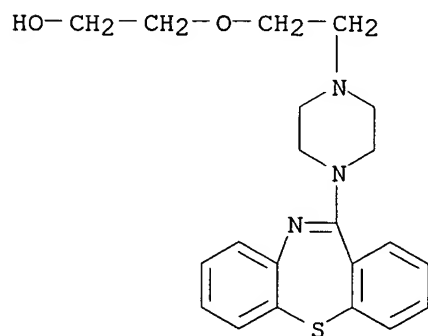
(9CI) (CA INDEX NAME)

HO-CH<sub>2</sub>-CH<sub>2</sub>-O-CH<sub>2</sub>-CH<sub>2</sub>



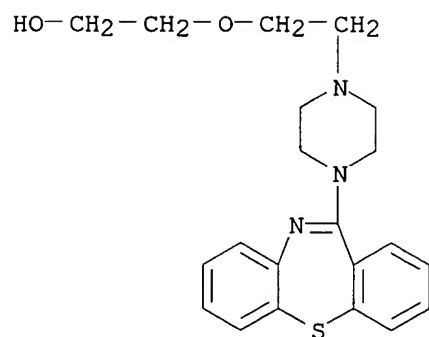
RE.CNT 5      THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS RECORD  
ALL CITATIONS AVAILABLE IN THE RE FORMAT

L30 ANSWER 39 OF 43 CAPLUS COPYRIGHT 2003 ACS  
 AN 1998:130310 CAPLUS  
 DN 128:225529  
 TI Focus on quetiapine: the fourth atypical antipsychotic  
 AU Caley, Charles F.; Rosenbaum, Susan  
 CS Burlingame Center for Psychiatric Research and Education, University of Connecticut, Institute of Living, Hartford, CT, USA  
 SO Formulary (1998), 33(2), 105-106, 109-110, 112, 115-116, 119  
 CODEN: FORMF9; ISSN: 1082-801X  
 PB Advanstar Communications, Inc.  
 DT Journal; General Review  
 LA English  
 AB A review with 30 refs. Quetiapine is a novel dibenzothiazepine-type atypical antipsychotic with moderate antagonism of dopamine type 1 and 2 and serotonin type 2a receptors. It is metabolized primarily by the CYP 3A4 isoenzyme and has poor estd. bioavailability (9%+-4%), relatively low protein binding (83%), and an elimination half-life of 6 h. Clin. trials show quetiapine to have favorable effects on the pos. and neg. symptoms of schizophrenia and to be more effective than placebo and as effective as chlorpromazine and haloperidol; direct comparisons with other atypical antipsychotics are unavailable. The drug's most frequent side effects are agitation, somnolence, headache, dry mouth, insomnia, postural hypotension, dizziness, and serum ALT elevations. Decreased serum thyroid hormone concns., elevated serum lipid levels, and wt. gain have also been reported, and the manufacturer warns of a risk of cataracts based on animal studies. Extrapyramidal reactions are infrequent with quetiapine, and the drug does not raise serum prolactin levels.  
 IT **111974-69-7**, Quetiapine  
 RL: ADV (Adverse effect, including toxicity); BPR (Biological process); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses)  
 (quetiapine as antipsychotic agent in humans)  
 RN 111974-69-7 CAPLUS  
 CN Ethanol, 2-[2-(4-dibenzo[b,f][1,4]thiazepin-11-yl-1-piperazinyl)ethoxy]-(9CI) (CA INDEX NAME)



L30 ANSWER 40 OF 43 CAPLUS COPYRIGHT 2003 ACS  
AN 1997:6767 CAPLUS  
DN 126:84470  
TI Seroquel restores sensorimotor gating in phencyclidine-treated rats  
AU Swerdlow, Neal R.; Bakshi, Vaishali; Geyer, Mark A.  
CS Dep. of Psychiatry and Neurosciences Program, Univ. of California, San Diego, La Jolla, CA, 2093-0804, USA  
SO Journal of Pharmacology and Experimental Therapeutics (1996), 279(3), 1290-1299  
CODEN: JPETAB; ISSN: 0022-3565  
PB Williams & Wilkins  
DT Journal  
LA English  
AB Phencyclidine (PCP) is a psychotomimetic noncompetitive glutamate antagonist that has been used in studies of the neural substrates of **psychosis**. Both schizophrenic patients and PCP-treated rats exhibit reduced amts. of prepulse inhibition (PPI) of the startle reflex, which is the normal inhibition of startle that occurs when the startling noise is preceded 30 to 500 ms by a weak prepulse. The present study assessed the effects of seroquel (ICI 204,636), a mixed D2/5-hydroxytryptamine2 antagonist with a preclin. profile suggestive of potential antipsychotic efficacy, on the PCP-induced disruption of PPI. Clozapine, risperidone and haloperidol were also studied as comparison compds. PCP (1.25 mg/kg) significantly reduced PPI, with prepulses that were 1 to 12 dB above background. Seroquel and clozapine significantly restored PPI in PCP-treated rats, whereas haloperidol and risperidone did not. Similar findings were obtained in studies using sep. animals, a slightly lower dose of PCP (1.0 mg/kg) and a high dose of each of these antipsychotics. Sep. studies verified that risperidone and haloperidol restored PPI in apomorphine-treated rats. In the present studies, seroquel exhibited a profile consistent with those exhibited by other "atypical" antipsychotics.  
IT 111974-72-2, Seroquel  
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(seroquel restores sensorimotor gating in phencyclidine-treated rats in relation to antipsychotic activity)  
RN 111974-72-2 CAPLUS  
CN Ethanol, 2-[2-(4-dibenzo[b,f][1,4]thiazepin-11-yl-1-piperazinyl)ethoxy]-, (2E)-2-butenedioate (2:1) (salt) (9CI) (CA INDEX NAME)  
CM 1  
CRN 111974-69-7  
CMF C21 H25 N3 O2 S

10/009,574

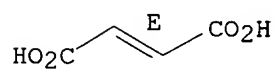


CM 2

CRN 110-17-8

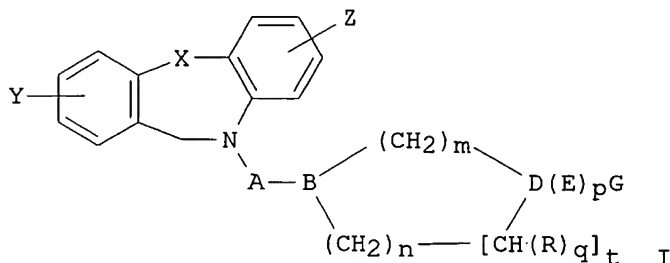
CMF C4 H4 O4

Double bond geometry as shown.



~~DB~~ ANSWER 41 OF 43 CAPLUS COPYRIGHT 2003 ACS  
~~AN~~ 1995:205963 CAPLUS  
 DN 123:9468  
 TI 2-, 3-, 4-, 5-, 6-, 7-, 8-, 9- and/or 10-substituted dibenzoxazepine and  
 dibenzthiazepine compounds as analgesics and prostaglandin E2 antagonists,  
 pharmaceutical compositions and methods of use  
 IN Hansen, Donald W., Jr.; Peterson, Karen B.  
 PA Searle, G. D., and Co., USA  
 SO U.S., 39 pp.  
 CODEN: USXXAM  
 DT Patent  
 LA English  
 FAN.CNT 3

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	US 5354747	A	19941011	US 1993-79021	19930616
	US 5461047	A	19951024	US 1994-245349	19940518
	WO 9429286	A1	19941222	WO 1994-US6029	19940602
	W:	AT, AU, BB, BG, BR, BY, CA, CH, CN, CZ, DE, DK, ES, FI, GB, HU, JP, KP, KR, KZ, LK, LU, LV, MG, MN, MW, NL, NO, NZ, PL, PT, RO, RU, SD, SE, SI, SK, TT, UA, US, UZ, VN			
	RW:	AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG			
	CA 2165159	AA	19941222	CA 1994-2165159	19940602
	AU 9471387	A1	19950103	AU 1994-71387	19940602
	EP 703908	A1	19960403	EP 1994-920687	19940602
	R:	AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LI, LU, NL, PT, SE			
	JP 09500107	T2	19970107	JP 1994-501874	19940602
PRAI	US 1993-79021		19930616		
	US 1994-245349		19940518		
	WO 1994-US6029		19940602		
OS	MARPAT 123:9468				
GI					



AB The present invention provides substituted dibenzoxazepine and  
 dibenzthiazepine compds. I which are useful as analgesic agents for the  
 treatment of pain, and for prostaglandin-E2 mediated diseases,  
 pharmaceutical compns. comprising a therapeutically-effective amt. of I in  
 combination with a pharmaceutically-acceptable carrier, a method for  
 eliminating or ameliorating pain in an animal comprising administering a  
 therapeutically-effective amt. of I to the animal, and a method for  
 treating prostaglandin-E2 mediated diseases in an animal comprising  
 administering a therapeutically-effective amt. of I to the animal.  
 Analgesic activity was measured using the writhing assay at std. dose of  
 10 mpk/g body wt.: I produced analgesia in from 2/10 to 10/10 of



the mice. Prostaglandin E2 antagonism assay (inhibition of contraction of guinea pig ileum): dose ratio of EC50 doses of from 0.8 to 32. Pharmaceutical compns. were given.

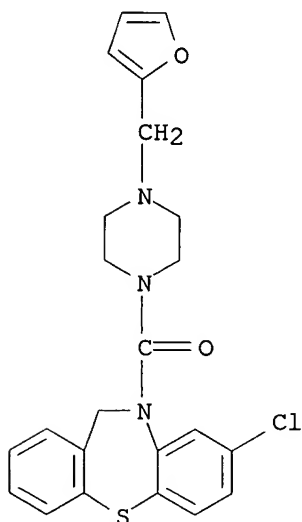
IT **163839-57-4P**, 1-[(8-Chlorodibenz[b,f][1,4]thiazepin-10(11H)-yl)carbonyl]-4-(2-furanylmethyl)piperazine **163839-58-5P**, 1-[(8-Chlorodibenz[b,f][1,4]thiazepin-10(11H)-yl)carbonyl]-4-(2-furanylmethyl)piperazine monohydrochloride

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(substituted dibenzoxazepine and dibenzthiazepine compds. as analgesics and prostaglandin E2 antagonists)

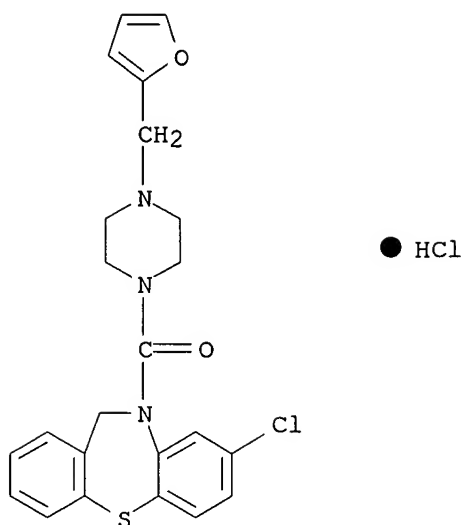
RN 163839-57-4 CAPLUS

CN Dibenzo[b,f][1,4]thiazepine, 8-chloro-10-[[4-(2-furanylmethyl)-1-piperazinyl]carbonyl]-10,11-dihydro- (9CI) (CA INDEX NAME)

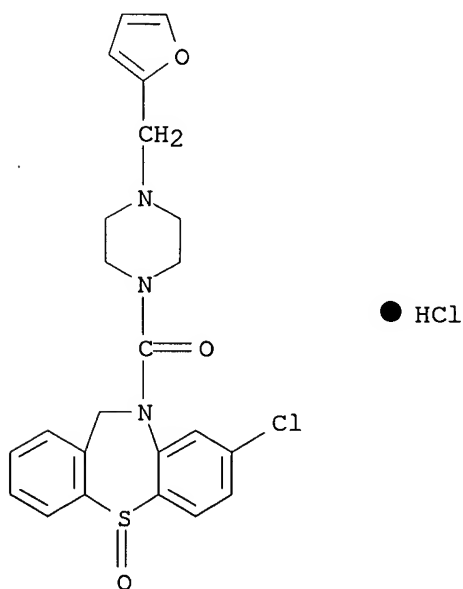


RN 163839-58-5 CAPLUS

CN Dibenzo[b,f][1,4]thiazepine, 8-chloro-10-[[4-(2-furanylmethyl)-1-piperazinyl]carbonyl]-10,11-dihydro-, monohydrochloride (9CI) (CA INDEX NAME)



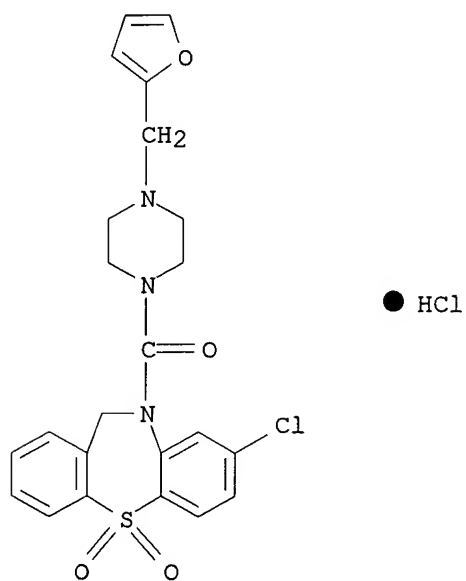
IT **163839-59-6P**, 1-8 (8-Chlorodibenz[b,f][1,4]thiazepin-10(11H)-yl)carbonyl]-4-(2-furanylmethyl)piperazine S-oxide monohydrochloride  
**163839-60-9P**, 1-[(8-Chlorodibenz[b,f][1,4]thiazepin-10(11H)-yl-carbonyl]-4-(2-furanylmethyl)piperazine S,S-dioxide monohydrochloride  
 RL: SPN (Synthetic preparation); PREP (Preparation)  
 (substituted dibenzoxazepine and dibenzthiazepine compds. as analgesics and prostaglandin E2 antagonists)  
 RN 163839-59-6 CAPLUS  
 CN Dibenzo[b,f][1,4]thiazepine, 8-chloro-10-[[4-(2-furanylmethyl)-1-piperazinyl]carbonyl]-10,11-dihydro-, 5-oxide, monohydrochloride (9CI)  
 (CA INDEX NAME)



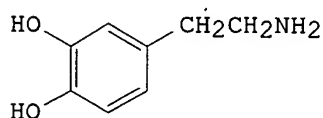
RN 163839-60-9 CAPLUS  
 CN Dibenzo[b,f][1,4]thiazepine, 8-chloro-10-[[4-(2-furanylmethyl)-1-

10/009,574

piperazinyl]carbonyl]-10,11-dihydro-, 5,5-dioxide, monohydrochloride (9CI)  
(CA INDEX NAME)



~~DS~~0 ANSWER 42 OF 43 CAPLUS COPYRIGHT 2003 ACS  
~~AN~~ 1976:99454 CAPLUS  
 DN 84:99454  
 TI Antagonism of the hyperactivity induced by dopamine applied  
 intracerebrally to the nucleus accumbens septi by typical neuroleptics and  
 by clozapine, sulpiride and thioridazine  
 AU Costall, Brenda; Naylor, Robert J.  
 CS Postgrad. Sch. Stud. Pharmacol., Univ. Bradford, Bradford/Yorkshire, UK  
 SO European Journal of Pharmacology (1976), 35(1), 161-8  
 CODEN: EJPHAZ; ISSN: 0014-2999  
 DT Journal  
 LA English  
 GI



AB Dopamine-HCl (I-HCl) [62-31-7] (50 .mu.g), administered intracerebrally to  
 the nucleus accumbens septi of rats, induced a dose-dependent  
 hyperactivity following pretreatment with nialamide. This I-induced  
 hyperactivity was inhibited by the i.p. injection of both typical  
 neuroleptic agents, haloperidol [52-86-8], pimozide [2062-78-4],  
 fluphenazine-HCl [146-56-5], and clothiapine [2058-52-8]  
 (0.05-0.5 mg/kg i.p.) and the atypical neuroleptics clozapine [5786-21-0],  
 sulpiride [15676-16-1] and thioridazine-HCl [130-61-0] (0.5-20 mg/kg i.p.)  
 although, generally, the doses required of the latter were in the order of  
 20-100 times those of the typical agents to produce an equiv. effect. In  
 contrast, cataleptic doses of metoclopramide-HCl [7232-21-5] (10-30 mg/kg  
 i.p.) failed to reduce the I-induced hyperactivity: aceperone [807-31-8]  
 and propranolol-HCl [318-98-9] were similarly ineffective. However,  
 inhibition of hyperactivity was recorded following the peripheral  
 administration of the antimanic drug, IB503 [14942-31-5]. Thus, the  
 ability of a drug to antagonize the hyperactivity induced by the injection  
 of I into the nucleus accumbens septi may be of value in the detection of  
 antipsychotic activity.

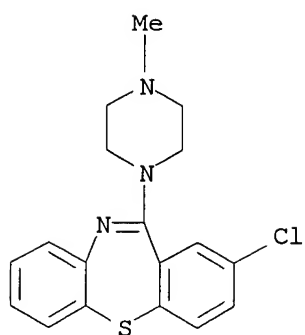
IT 2058-52-8

RL: BIOL (Biological study)  
 (hyperactivity from dopamine response to, antipsychotic activity in  
 relation to)

RN 2058-52-8 CAPLUS

CN Dibenzo[b,f][1,4]thiazepine, 2-chloro-11-(4-methyl-1-piperazinyl)- (7CI,  
 8CI, 9CI) (CA INDEX NAME)

10/009,574



~~LSO~~ ANSWER 43 OF 43 CAPLUS COPYRIGHT 2003 ACS  
~~AN~~ 1973:427467 CAPLUS  
~~DN~~ 79:27467  
TI Toxicity studies with metiapine  
AU Gibson, J. P.; Rohovsky, M. W.; Newberne, J. W.; Larson, E. J.  
CS Megrell-Natl. Lab. Div., Richardson-Merrell Inc., Cincinnati, OH, USA  
SO Toxicology and Applied Pharmacology (1973), 25(2), 220-9  
CODEN: TXAPA9; ISSN: 0041-008X  
DT Journal  
LA English  
AB Continuous daily dietary administration of 3, 10, and 30 mg/kg doses of metiapine (I) [5800-19-1] to rats for 18 months produced a dose-related degree of depression and decreased food consumption and body wt. gain. The acute oral LD50 in mice and rats was 680 and 943 mg/kg, resp. Dogs showed varying degrees of depression and stimulation when given single daily oral doses of 5, 15, or 50 mg/kg for 1 year, and mammary enlargement with milk prodn. was obsd, in some of the females. The 50 mg/kg/day dogs showed slight increases in serum alk. phosphatase [9001-78-9] activity. Except for the mild alk.phosphatase changes, the effects obsd, were attributed to the psychotropic activity of I, and its secondary effects on appetite and endocrine function.  
IT **5800-19-1**  
RL: ADV (Adverse effect, including toxicity); BIOL (Biological study) (toxicity of)  
RN 5800-19-1 CAPLUS  
CN Dibenzo[b,f][1,4]thiazepine, 2-methyl-11-(4-methyl-1-piperazinyl)- (7CI, 8CI, 9CI) (CA INDEX NAME)

